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(54) Resurfacing of rodent antibodies.

67 A method for determining how to humanize a rodent antibody or fragment thereof by resurfacing, said method comprising:

(a) determining the conformational structure of the variable region of said rodent antibody or fragment thereof by constructing a three-dimensional model of said rodent antibody variable region;

(b) generating sequence alignments from relative accessibility distributions from x-ray crystallographic structures of a sufficient number of rodent antibody variable region heavy and light chains to give a set of heavy and light chain framework positions wherein said set is identical in 98% of said sufficient number of rodent antibody heavy and light chains;

(c) defining for said rodent antibody or fragment thereof to be humanized a set of heavy and light chain surface exposed amino acid residues using said set of framework positions generated in said step

(b);

(d) identifying from human antibody amino acid sequences a set of heavy and light chain surface exposed amino acid residues that is most closely identical to said set of surface exposed amino acid residues defined in said step (c), wherein said heavy and light chain from said human antibody are or are not naturally paired;

(e) substituting, in the amino acid sequence of said rodent antibody or fragment thereof to be humanized said set of heavy and light chain surface exposed amino acid residues defined in said step (c) with said set of heavy and light chain surface exposed amino acid residues identified in said step (d);

(f) constructing a three-dimensional model of said variable region of said rodent antibody or fragment

thereof resulting from the substituting specified in said step (e);

(g) identifying, by comparing said three-dimensional models constructed in said steps (a) and (f), any amino acid residues from said set identified in said step (d), that are within 5 Angstroms of any atom of any residue of the complementarity determining regions of said rodent antibody or fragment thereof to be humanized; and

(h) changing any residues identified in said step (g) from the human to the original rodent amino acid residue to thereby define a rodent antibody humanizing set of surface xposed amino acid residues; with the provise that said step (a) need not be conducted first, but must be conducted prior to said step (g).

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#### FIELD OF THE INVENTION

The present invention relates to the development of prediction rules that can be used to accurately model the variable regions (V-regions) of antibodies. The development of these rules and their application in the predictive molecular restructuring of the surfaces of variable domains of non-human monoclonal antibodies enables changing of the surface, resurfacing, of these monoclonal antibody V-regions to replicate the surface characteristics found on human antibody V-regions. This method of resurfacing non-human monoclonal antibody V-regions to resemble human antibody V-regions is expected to permit the production of functional altered antibodies, which retain the binding parameters of the original non-human monoclonal antibody, with improved therapeutic efficacy in patients due to the presentation of a human surface on the V-region.

# **BACKGROUND OF THE INVENTION**

#### **General Background of Antibodies**

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Murine monoclonal antibodies are widely used as diagnostic and therapeutic agents in the treatment of human disease. Mice can be readily immunized with foreign antigens to produce a broad spectrum of high affinity antibodies. Invariably, the introduction of murine or other rodent antibodies into humans results in the production of a human anti-mouse antibody (HAMA) response due to the presentation of a foreign protein in the body. The production of HAMA in patients can result from the introduction of foreign antibody in a single dose or from extended use in therapy, for example, for the treatment of cancer. Extended use of murine antibody is generally limited to a term of days or weeks in patients before concerns of anaphylaxis arise. Moreover, once HAMA has developed in a patient, future use of murine antibodies for diagnostic or therapeutic purposes is often precluded for the same reasons.

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Beyond ethical considerations, attempts to produce human monoclonal antibodies have not been highly successful for a number of reasons. The production *in vitro* of human monoclonals rarely results in high affinity antibodies. *In vitro* cultures of human lymphocytes yield a restricted range of antibody responses relative to the broad spectrum of reactive antibodies produced *in vivo* through direct immunization of mice. Additionally, in humans, immune tolerance prevents the successful generation of antibodies to self-antigens. All of these factors have contributed to the search for ways to modify the structures of murine monoclonal antibodies to improve their use in patients. Many investigators have attempted to alter, reshape or humanize murine monoclonal antibodies in an effort to improve the therapeutic application of these molecules in patients.

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# Strategies of Antibody Humanization

The earliest reports of the controlled rearrangement of antibody domains to create novel proteins was demonstrated using rabbit and human antibodies as described by Bobrzecka, K. et al. (Bobrzecka, K., Konieczny, L., Laidler, P. and Rybarska, J. (1980), Immunology Letters 2, pp. 151-155) and by Konieczny et al. (Konieczny, L., Bobrzecka, K., Laidler, P. and Rybarska, J. (1981), Haematologia 14 (I), pp. 95-99). In those reports, the protein subunits of antibodies, rabbit Fab fragments and human Fc fragments, were joined through protein disulfide bonds to form new, artificial protein molecules or chimeric antibodies.

Recombinant DNA technology was used to construct gene fusions between DNA sequences encoding mouse antibody variable light and heavy chain domains and human antibody light chain (LC) and heavy chain (HC) constant domains to permit expression of the first recombinant "near-human" antibody (chimeric antibody) product (Morrison, S.L., Johnson, M.J., Herzenberg, L.A. and Oi, V.T. (1984), Proc. Natl. Acad. Sci. U.S.A. 81, pp. 6851-6855).

The kinetics and immune response in man to chimeric antibodies has been examined (LoBuglio, A.F., Wheeler, R.H., Trang, J., Haynes, A., Rogers, K., Harvey, E.B., Sun, L., Ghrayeb, J. and Khazaeli, M.B. (1989), Proc. Natl. Acad. Sci. **86**, pp. 4220-4224).

Chimeric antibodies contain a large number of non-human amino acid sequences and are immunogenic in man. The result is the production of human anti-chimera antibodies (HACA) in patients. HACA is directed against the murine V-region and can also be directed against the novel V-region/C-region (constant region) junctions present in recombinant chimeric antibodies.

To overcome some of the limitations presented by the immunogenicity of chimeric antibodies, the DNA sequences encoding the antigen binding portions or complementarity determining regions (CDR's) of murine monoclonal antibodies have been grafted by molecular means in the DNA sequences encoding the frameworks of human antibody heavy and light chains (Jones, P.T., Dear, P.H., Foote, J., Neuberger, M.S. and Winter, G. (1986), Nature 321, pp. 522-525; Riechmann, L., Clark, M., Waldmann, H. and Winter, G. (1988), Nature 332,

pp. 323-327). The expressed recombinant products called reshaped or humanized antibodies are comprised of the framework of a human antibody light or heavy chain and the antigen recognition portions, CDR's, of a murine monoclonal antibody. Several patent applications have been filed in this area including, for example, European Patent Application, Publication No. 0239400; European Patent Application, Publication Nos. 0438310A1 and 0438310A2; International Patent Publication No. WO 91/09967; and International Patent Publication No. WO 90/07861.

However, it is questionable whether European Patent Application (EP), Publication No. 0239400 is truly enabling. It is not assured in this patent that the best fit is made to assure proper presentation of the CDR loops at the antibody combining site.

EP Publication Nos. 0438310A1 and 0438310A2 go a step beyond EP Publication No. 0239400 by protecting the importance of uniquely selected human frameworks for the human light chain (LC) and heavy chain (HC) V-regions. These V-region frameworks should show a high degree of sequence similarity with the frameworks of the murine monoclonal antibody and present the CDR's in the appropriate configuration. However, the criteria for sequence matching are no more sophisticated than simple homology searching of the antibody protein or DNA databases.

International Patent Publication No. WO 91/09967 attempts a further variation of the method disclosed in EP Publication No. 0239400. In International Patent Publication No. WO 91/09967, homology of the donor sequences and the acceptor framework is not important, rather it discloses that a selected set of residues in the LC and HC are critically important to humanization. The ability to make changes at these positions is the basis of International Patent Publication No. WO 91/09967.

International Patent Publication No. WO 90/07861 proposes four important criteria for designing humanized antibodies. 1) Homology between human acceptor and non-human donor sequences. 2) Use donor rather than acceptor amino acids where the acceptor amino acid is unusual at that position. 3) Use donor framework amino acids at positions adjacent to the CDR. 4) Use donor amino acids at framework positions where the sidechain atom is within 3 Angstroms of the CDR in a 3-D model. The first antibody humanized by this method retained less than 1/3 the affinity of the original monoclonal antibody.

None of the above methods for designing a humanized antibody are predictable due to the questions that surround CDR framework interactions. By replacement of murine framework with human framework, there is no guarantee of identical conformations for CDR's because i) the V<sub>L</sub>-V<sub>H</sub> interaction is not identical in all V-regions and ii) accurate prediction of the CDR-framework interactions are key to faithful reproduction of the antigen binding contacts.

The above methods do not offer a general solution to solving the issues surrounding antibody humanization, rather the methods as outlined in each reference above involve a substantial amount of trial and error searching to obtain the desired affinity in the final humanized product. More importantly, there is no guarantee that corrective changes in framework amino acids will leave the reshaped V-regions resembling the surface character of a truly human antibody. Therefore, it can be argued that antibodies humanized by the above methods may be immunogenic in man.

#### **Antigenicity of Antibodies**

The antigenicity/immunogenicity of an antibody, including recombinant reshaped antibody products, introduced into humans can be viewed as a surface phenomenon. In general one can view the immune system as scanning the surface of a protein introduced to the body. If the  $F_V$  portion of a humanized antibody 'opensup' in the circulation then internal residues can be presented to the immune system. On the other hand, if the  $F_V$  portion is stable and tightly packed then only the surface residues presented by the V-regions and the interface between the  $V_L$  and  $V_H$  regions will be 'scanned'.

# Surface Reshaping or Resurfacing of Antibobies

The notion of surface presentation of proteins to the immune system raises the prospect of redesigning murine monoclonal antibodies to resemble human antibodies by humanizing only those amino acids that are accessible at the surface of the V-regions of the recombinant  $F_V$ . The resurfacing of murine monoclonal antibodies to reduce their immunogenicity could be beneficial in maintaining the avidity of the original monoclonal antibody in the reshaped version, because the natural framework-CDR interactions are retained. The value of maintaining the integrity of the framework-CDR interactions has been illustrated as summarized below.

In a rec nt research report, two different reshaped versions of the rat monoclonal antibody, Campath-9 (anti-human CD4), were generated (Gorman, S.D., Clark, M.R., Routledge, E.G., Cobbold, S.P. and Waldmann, H. (1991), Proc. Natl. Acad. Sci. U.S.A. 88, pp. 4181-4185). In one version, pV<sub>H</sub>NEW/C<sub>G1</sub>, the acceptor V<sub>H</sub> fra-

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mework was from the human NEW-based heavy chain, which has 47% identical residues to the Campath-9  $V_H$ . While in the second version,  $pV_HKOL/C_{G1}$ , the acceptor  $V_H$  framework was from the human KOL antibody, which has 72% identical residues to Campath-9  $V_H$ . Each reshaped antibody contained the identical  $V_L$  domain from the human REI antibody sequence. However, the recombinant product of  $pV_HKOL/C_{G1}$  had an avidity for CD4 that was substantially greater than the product of  $pV_HNEW/C_{G1}$ . The authors proposed a reshaping strategy where human sequences, that are highly homologous to the rodent antibody of interest, are transferred, by in vitro mutagenesis, into the rodent V-region to create a "bestfit" reshaped antibody. This strategy uses the term "bestfit" to describe the modeling process, however, there is no quantitative formula employed to assess "bestfit", and so in effect, the process is subjective. Additionally, there is no resurfacing concept presented in that paper.

The concept of reducing rodent-derived antibody immunogenicity through the replacement of exposed residues in the antibody framework regions which differ from those of human origin is discussed in a recent paper (Padlan, E.A. (1991), Molecular Immunology 28, pp. 489-498). In that paper, the variable domains of two antibody structures, KOL (human) and J539 (mouse), are examined. The crystal structures of the Fab fragments of these two antibodies have been elucidated to high resolution. The solvent accessibility of the exposed framework residues in the variable domains of these two antibodies were compared to a sequence database of human and murine antibody V-region subgroups. On the basis of his findings, Padlan proposed to reduce the antigenicity of allogeneic variable domains [murine V-regions], through replacement of the exposed residues in the framework regions with residues usually found in human antibodies. In murine sequences with the highest similarity to a given human sequence, the number of changes necessary to "humanize" a murine V-region surface would range from 6-15 amino acid changes per V-region. This reference suggests how to convert one antibody surface into another but no general method is developed. Application of the procedure is provided by two examples, a worst-case and a best-case.

#### Worst Case:

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Among the representative murine kappa  $V_L$  sequences examined for which its autologous  $V_H$  has b n sequenced, S107V<sub>L</sub> has the most residues that need to be replaced to humanize it. S107V<sub>L</sub> is most similar to the members of the human subgroup VKIV and JK2. The exposed or partially exposed residues that need to be replaced are those at positions  $\underline{9}$ , 10, 14,  $\underline{15}$ , 16, 17, 18,  $\underline{22}$ ,  $\underline{41}$ ,  $\underline{63}$ , 80,  $\underline{83}$ ,  $\underline{85}$ , 100 and 106. Murine V-region S107V<sub>H</sub> is most similar in its framework to the members of the human subgroup VHIII and JH6. The exposed or partially exposed residues in S107V<sub>H</sub> that need to be replaced are those at positions 3, 40, 68, 73, 75, 76, 82b and 89. A total of 23 residues need to be replaced to humanize the variable domains of S107.

#### Best Case:

Among the murine  $V_H$  sequences examined for which the autologous  $V_L$  has also been sequenc d, MOPC21V<sub>H</sub> has the least number of residues that need to be replaced to humanize it. MOPC21V<sub>H</sub> is most similar in its framework to the members of the human subgroup HIII and JH6. The exposed or partially exposed residues that need to be replaced are those at positions 1, 42, 74, 82a, 84, 89 and 108. MOPC21V<sub>L</sub> is most similar in its framework to human subgroup VKIV and JK4. The exposed or partially exposed residues that need to be replaced are those at positions 1, 9, 12, 15, 22, 41, 63, 68, 83 and 85. A total of 17 amino acids need to be replaced to humanize the variable domains of MOPC21.

Of the light chains in the Best- and Worst-Case examples cited above, S107V<sub>L</sub> required changes at 15 positions and MOPC21V<sub>L</sub> required changes at 10 positions. Only seven of the changes are common to both of these light chain sequences (see underlined residues). Moreover, of the heavy chain residues that need to be replaced to humanize the respective V-regions, S107V<sub>H</sub> required changes at 8 positions and MOPC21V<sub>H</sub> required changes at 7 positions. In this instance, only one position is common to both of these heavy chain sequences (see residues in boldface).

An analysis of S107 V-regions alone would not have led to the prediction of which residues to change in MOPC21. The reason for this is that the surface residues in Padlan's analysis are only determined by reference to the crystal structure analysis of <u>one</u> antibody. In addition, the basis for defining the surface exposure of an amino acid at a particular position on that crystal structure is a continuous gradient of change, e.g., the fractional solvent accessibility values (Padlan, E.A. (1990), Molecular Immunology 28, pp. 489-498) were computed, where: 0 to 0.2 = completely buried, 0.2 to 0.4 = mostly buried, 0.4 to 0.6 = partly buried/partly exposed, 0.6 to 0.8 = mostly expos d, and 0.8 or above = completely exposed. By limiting the analysis of exposed surface residues to a single crystal structure and by superimposing a broad range of solvent accessibility ratios on exposed residues, such a modeling strategy could be expected to have a wide margin of error in its calculations.

This model fails to take into account the great majority of structural information available in the database for other antibody crystal structures.

#### SUMMARY OF THE INVENTION

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Accordingly, it is an object of this invention to provide humanized rodent antibodies or fragments thereof, and in particular, humanized rodent monoclonal antibodies that have improved therapeutic efficacy in patients due to the presentation of a human surface on the V-region. This and other objects have been attained by providing a method for determining how to humanize a rodent antibody or fragment thereof by resurfacing the method comprising:

- (a) determining the conformational structure of the variable region of the rodent antibody or fragment thereof by constructing a three-dimensional model of the rodent antibody variable region;
- (b) generating sequence alignments from relative accessibility distributions from x-ray crystallographic structures of a sufficient number of rodent antibody variable region heavy and light chains to give a set of heavy and light chain framework positions wherein the set is identical in 98% of the sufficient number of rodent antibody heavy and light chains;
- (c) defining for the rodent antibody or fragment thereof to be humanized a set of heavy and light chain surface exposed amino acid residues using the set of framework positions generated in step (b);
- (d) identifying from human antibody amino acid sequences a set of heavy and light chain surface exposed amino acid residues that is most closely identical to the set of surface exposed amino acid residues defined in step (c), wherein the heavy and light chain from the human antibody are or are not naturally paired;
- (e) substituting, in the amino acid sequence of the rodent antibody or fragment thereof to be humanized the set of heavy and light chain surface exposed amino acid residues defined in step (c) with the set of heavy and light chain surface exposed amino acid residues identified in step (d);
- (f) constructing a three-dimensional model of the variable region of the rodent antibody or fragment thereof resulting from the substituting specified in step (e);
- (g) identifying, by comparing the three-dimensional models constructed in steps (a) and (f), any amino acid residues from the set identified in step (d), that are within 5 Angstroms of any atom of any residue of th complementarity determining regions of the rodent antibody or fragment thereof to be humanized; and
- (h) changing any residues identified in step (g) from the human to the original rodent amino acid residu to thereby define a rodent antibody humanizing set of surface exposed amino acid residues; with the proviso that step (a) need not be conducted first, but must be conducted prior to step (g).

Also provided is a method for producing a humanized rodent antibody or fragment thereof from a rod nt antibody or fragment thereof, the method comprising:

- (I) carrying out the above-described method for determining how to humanize a rodent antibody or fragment thereof by resurfacing; and
- (II) modifying the rodent antibody or fragment thereof by replacing the set of rodent antibody surface x-posed amino acid residues with the rodent antibody humanizing set of surface exposed amino acid residues defined in step (h) of the above-described method.
- In a preferred embodiment, the rodent antibody or fragment thereof is a murine antibody, and most pr ferably murine antibody N901.

# BRIEF DESCRIPTION OF THE FIGURES

Figure 1 shows an algorithm that can be used for constructing a three-dimensional model of the rodent antibody variable region.

Figure 2 is a diagram showing the approach to determine how to humanize a rodent antibody or fragment thereof according to the present invention.

Figures 3A and 3B are plots of relative accessibility of amino acid residues for twelve antibody F<sub>V</sub> structures, mapped onto the sequence alignment of these structures. Structures Glb2 (Jeffrey, P.D., Doctor of Philosophy Thesis, University of Oxford, United Kingdom, 1991), D1.3 (Amit, A.G., Mariuzza, R.A., Phillips, S.E.V. and Poljak, R.J. (1986), Science 233, pp. 747-753), 3D6 (Grunow, R., Jahn, S., Porstman, T., Kiessig, T., Steinkeller, H., Steindl, F., Mattanovich, D., Gurtl r, L., Deinhardt, F., Kating r, H. and von R., B. (1988), J. Immunol. Meth. 106, pp. 257-265) and 36-71 (5fab) (Rose, D.R., Strong, R.K., Margolis, M.N., Gefter, M.L. and Petsko, G.A. (1990), Proc. Natl. Acad. Sci. U.S.A. 87, pp. 338-342) ar not y t present in th Brookhaven database. The other structures used wer: 2hfl (Sheriff, S., Silverton, E.W., Padlan, E.A., Cohen, G.H., Smith-Gill, S.J., Finzel, B.C. and Davies, D.R. (1987), Proc. Natl. Acad. Sci. U.S.A. 84, pp. 8075-8079), 3hfm (Padlan, E., Silverton, E., Sheriff, S., Cohen, G., Smith-Gill, S. and Davies, D. (1989), Proc. Natl. Acad. Sci. U.S.A. 86, pp.

5938-5942), 2fbj (Mainhart, C.R., Potter, M. and Feldmann, R.J. (1984), Mol. Immunol. 21, pp. 469-478), 3fab (Saul, F.A., Amzel, L.M. and Poljak, R.J. (1978), J. Biol. Chem. 253, pp. 585-597), 4fab (Herron, J., He, X., Mason, M., Voss, E. and Edmunson, A. (1989), Proteins: Struct., Funct., Genet. 5, pp. 271-280), 2mcp (Segal, D., Padlan, E., Cohen, G., Rudikoff, S., Potter, M. and Davies, D. (1974), Proc. Natl. Acad. Sci. U.S.A. 71, pp. 4298-[??]), 2fb4 (Marquart, M. Deisenhofer, J. and Huber, R. (1980), J. Mol. Biol. 141, pp. 369-391), and 1f19 (Lascombe, M. Alzari, P., Boulot, G., Salujian, P., Tougard, P., Berek, C., Haba, S., Rosen, E., Nisonof, A. and Poljak, R. (1989), Proc. Natl. Acad. Sci. U.S.A. 86, p. 607). These structures are designated by their Brookhaven entry code. The sequence numbering used here is described in Figures 4A and 4B. Figure 3A graphically shows the relative accessibility for the heavy chain and Figure 3B graphically shows the relative accessibility for the light chain.

Figures 4A and 4B show alignments of sequences generated using the three methods of humanization. Sequences are: 1) Original rodent N901. 2+3) KOL (Marquart, M. Deisenhofer, J. and Huber, R. (1980), J. Mol. Biol. 141, pp. 369-391) and reshaped N901 using KOL surface. 4+5) Most homologous sequences, L(KV2F) (Klobeck, H., Meindl, A., Combriato, G., Solomon, A. and Zachau, H. (1985), Nucleic Acids Res. pp. 6499-6513) and H(G36005) (Schroeder, H. and Wang, J. (1990), Proc. Natl. Acad. Sci. U.S.A. 87), and reshaped N901 using these sequences. 6+7) Most homologous with respect to surface residues, L(KV4B) (Klobeck, H., Bronkamp, G., Combriato, G., Mocikat, R., Pohelnz, H. and Zachau, H. (1985), Nucleic Acids Res. 3, pp. 6515-6529) and H(PLO123) (Bird, J., Galili, N., Link, M., Sites, D. and Sklar, J. (1988), J. Exp. Med. 168, pp. 229-245), and reshaped N901 using these sequences. The numbering is the same as used in the antibody modelling program ABM (trademark for commercial software, Oxford Molecular Ltd., Oxford, U.K.), which is based on structural conservation and not sequence homology as used by Padlan et al. (Kabat, E.A., Wu, T.T., Reid-Miller, M., Perry, H.M. and Gottesman, K.S. (1987), Sequences of Proteins of Immunological Interest. U.S. Department of Health and Human Services, Fourth Edition). The sequence changes which have to be introduced in order to resurface N901 with a given sequence are marked with bars, back-mutations as determined from F<sub>V</sub> models are marked with stars. The sequence homology of given sequences to N901 are shown in brackets after each sequence

Figure 5 is a stereo plot of mean antibody  $\beta$ -barrel, coordinates determined by iterative multiple fitting of eight antibody structures. Strands 7 and 8 comprise the 'take off' positions for CDR H3 and are not included in the fitting of  $V_L$  and  $V_H$  regions.

Figure 6 is a plot of RMS deviation from the mean of the eight  $\beta$ -sheet strands comprising the framework. The RMS was calculated from structures F19.9, 4-4-20, NEW, FBJ, KOL, HyHEL-5, HyHEL-10 and McPC603. N,C $\alpha$ ,C atoms are included in the plot. The residues used are shown in the alignment (Table 2). The most disordered residues are all the residues of strand HFR4, the last residue of LFR1, and the first and last residue of HFR2. The nomenclature of the strands is explained in the alignment in Table 2. LFR1 - #1, LFR2 - #2, LFR3 - #3, LFR4 - #4, HFR1 - #5, HFR2 - #6, HFR3 - #7, HFRS4 - #8.

Figure 7 is a flowchart of the overall modelling protocol known as CAMAL.

Figure 8 is a plot of superimposed loop backbones for models and x-ray structures discussed in Example 2. The loops are positioned after global framework fit. This does not represent the best local least squares fit, but shows how the loops are positioned globally onto the framework.

Figures 9A to 9D are stereo (N,C- $\alpha$ ,C,O) representations of crystal structures and models of D1.3, 3671 and Gloop-2 variable domain and  $\beta$ -barrel strands described in Example 2. Crystal structures are shown with open bonds, model with solid bonds. The difference between the 3D6-H3 in the model and the crystal structure is due to a 5-7° twist in the extended  $\beta$ -sheet conformation of this loop, Figure 9A: D1.3, Figure 9B: 36-71, Figure 9C: Gloop-2, Figure 9D: 3D6.

Figure 10 is a histogram showing the distribution of loop length for CDR H3 loops, data from Kabat et al. (Kabat, E.A., Wu, T.T., Reid-Miller, M., Perry, H.M. and Gottesman, K.S. (1987), Sequences of Proteins of Immunological Interest. U.S. Department of Health and Human Services, Fourth Edition).

#### **DETAILED DESCRIPTION OF THE INVENTION**

The existence of specific, yet different, surface patches in murine and human antibodies may be the origin of the inherited immunogenicity of murine antibodies in humans. Statistical analysis of a database of uniqu human and murine antibody  $F_V$  fragments has revealed that certain combinations of residues in exposed surface positions are specific for human and murine sequences. The combinations are not the same in human and murine  $F_V$  domains. However, it is possible to define families of surface residues for the two species of antibodies. These families reveal a novel method for the "humanization" or reshaping of murine antibodies. Humanization is the modification of the solvent accessible surface of a non-human antibody or fragment thereof to resemble the surface of a chosen human antibody or fragment thereof such that the modified non-human antibody or fragment thereof exhibits low remnunogenicity when administered to humans. Such a process

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applies in the present application to antibody variable regions but could equally well apply to any other antibody fragment. The method is considered to be generally applicable to humanization of rodent antibodies.

According to the present invention, a statistical analysis is presented which is based on accessibility calculated for a range of antibody crystal structures. When this information is applied to an antibody sequence database, it is possible to discriminate between human and murine antibodies at the sequence level purely on the basis of their surface residue profiles.

# Rational Resurfacing Approach

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There are several key features of the resurfacing approach of the present invention.

- 1) This method uses as a starting point, construction of a three-dimensional model of a rodent variable region by known methods;
- 2) A large number (e.g., twelve) of antibody  $F_V$  or Fab fragment x-ray crystallographic structures are analyzed to produce an unambiguous set of surface exposed amino acid residues that will be positionally identical for a majority (98%) of antibodies. The set is produced by identifying all those residues whose solvent accessibility is above a given cut-off (typically 30%), calculated using a modification of the method of Kabsch and Sander (Kabsch, W. and Sander, C. (1983), Biopolymers 22, pp. 2257-2637) in which explicit atomic radii are used for each atom type to predict sidechain positions as is described below in more detail; 3) Using a complete human antibody database, the best set of human heavy and light chain surface exposed amino acid residues is selected on the basis of their closest identity to the set of surface amino acid residues of the murine antibody;
- 4) In order to retain the conformational structure- of the CDRs of the rodent antibody, replacement of any human surface exposed amino acid with the original rodent surface exposed amino acid residue is carried out whenever a surface residue is calculated from the three-dimensional model to be within 5 Angstroms of a CDR residues.

The general resurfacing approach of the present invention is illustrated in Figure 2. The approach can be divided into two stages. In the first, the rodent framework (white) is retained and only the surface residues changed from rodent (dark grey circles) to the closest human pattern (light grey circles). This should remove the antigenicity of the rodent antibody. In the second stage, surface residues within 5 Angstroms of the CDRs are replaced with the rodent equivalents in an attempt to retain antigen binding and CDR conformation.

The method of the present invention is applicable to whole antibodies as well as antibody fragments. Suitable antibody fragments that can be used can readily be determined by the skilled artisan. Examples of some suitable fragments include a single chain antibody (SCA), an antibody  $F_V$  fragment, Fab fragment, Fab fragment, Fab' fragment, or other portion of an antibody comprising the binding site thereof.

According to the present invention, an important step in the method for determining how to modify a rodent antibody or fragment thereof by resurfacing is to determine the conformational structure of the variable region of the rodent antibody or fragment thereof to be humanized by constructing a three-dimensional model of the rodent antibody variable region. This can be done by known methods such as those described, for example, in Martin et al. (Martin, A.C.R., Cheetham, J.C. and Rees, A.R. (1989), Proc. Natl. Acad. Sci. U.S.A. 86, pp. 9268-9272; Methods in Enzymology (1991), 203, pp. 121-152) and as described in detail in Example 2.

Martin et al. describe an algorithm which is depicted in Figure 1. The algorithm applies to murine and human antibodies equally well. The present inventors therefore expect that, based on sequence similarity between antibodies of different species (Kabat, E.A. Segments of Proteins of Immunological Interest, National Institutes of Health, U.S.A. 1991), the algorithm will work equally well for rat and other rodent antibodies.

Briefly, the algorithm depicted in Figure 1 can be summarized as follows. The framework region of an antibody to be modelled is selected on the basis of sequence homology and constructed by a least squares fit onto the six conserved strands of the variable region β-barrel. Light and heavy chain complementarity determining regions are constructed using a combination of canonical structures (Chothia, C. and Lesk, A.M. (1987), J. Molec. Bio. 196, pp. 901-917), database searching and conformational searching. Detailed descriptions of these methods are described in Example 2 herein and in the above two references (Martin et al. 1989 and 1991).

According to the present invention, another three-dimensional model is also constructed. The other three-dimensional model is of the rodent antibody variable region having human antibody surface amino acid residues substituted therein at particular rodent antibody surface residue positions.

This other thr e-dimensional model is constructed by carrying out the series of steps described next.

The first of the steps is to generate sequence alignments from relative accessibility distributions from x-ray crystallographic structures of a sufficient number of rodent antibody variable region heavy and light chains to give a set of framework positions of surface xposed amino acid residues which is identical in a majority

(98%) of the variable regions.

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As used herein, the term "framework" means the antibody variable region from which the complementarity determining regions have been excluded.

"Complementarity determining regions" means those amino acid sequences corresponding to the following numbering system as defined by Kabat, E.A. (In Sequences of Immunological Interest, N.I.H., U.S.A., 1991).

Light Chain	L1	residues	24-34
Light Chain	L2	residues	50-56
Light Chain	L3	residues	89-97
Heavy Chain	H1	residues	31-358
Heavy Chain	H2	residues	50-58
Heavy Chain	Н3	residues	95-102

A sufficient number of rodent antibody fragments that need to be analyzed in order to produce the set of framework positions of surface exposed amino acid residues can readily be determined by the skilled artisan through routine experimentation using a database of antibody sequences. Thus, this step can be conducted using suitable databases now in existence or later compiled.

The x-ray crystallographic structures are used to determine relative accessibility distributions of surface exposed amino acid residues. The relative accessibility distributions identify all those residues whose solvent accessibility is above a given cut-off (typically 30%), calculated using a modification of the method of Kabsch and Sander (Kabsch, W. and Sander C. (1983), Biopolymers 22, pp. 2257-2637) in which explicit atomic radii are used for each atom type.

The relative accessibility distributions determined from the x-ray crystallographic structures can then be used to generate sequence alignments which give a set of framework positions of surface exposed amino acid residues which is identical in a majority (98%) of the variable regions.

The set of framework positions of surface exposed amino acid residues for the variable regions of murine antibodies is shown in Table 1, set forth in Example 1, and was produced using the sequence alignments and accessibility distributions shown in Figures 3A and 3B.

Once a set of framework positions of surface exposed amino acid residues for the variable regions of the rodent antibodies have been generated, the surface exposed residues of the heavy and light chain pair of the rodent antibody, or fragment thereof, to be humanized can be identified using an alignment procedure such as that described in Example 1 and shown in Figures 3A and 3B. This defines a set of surface exposed amino acid residues of a heavy and light chain pair of a rodent antibody or antibody fragment to be humanized.

Next, a complete human antibody sequence database is used to identify a set of surface exposed amino acid residues from a human antibody variable region that have the closest positional identity to the set of surface exposed amino acid residues of the variable region of the rodent antibody that is to be humanized. The set of surface exposed amino acid residues from the human antibodies can be separately identified for a heavy chain and for a light chain that are not naturally paired and/or a set can be identified from a natural human heavy and light chain pair, that is, a pair originating from a single B cell or hybridoma clone. Preferably, th set is one from a natural human heavy and light chain pair.

A humanized rodent antibody that gives the appearance of a human antibody is then predicted by substituting the set of surface exposed amino acid residues from the rodent antibody or fragment thereof to be humanized with the set of surface exposed amino acid residues from the human antibody.

A three-dimensional model can then be constructed from the resulting, fully substituted variable region of the rodent antibody or fragment thereof. The three-dimensional model is constructed using the same known methods mentioned above for constructing a 3-D model of the original rodent antibody or fragment there f.

While the antigenicity of this fully "resurfaced" or humanized antibody should be removed, an additional factor to be addressed is the binding affinity or the binding strength of the resurfaced antibody. Changes in the fram work of the variable domain introduced through resurfacing can influence the conformation of the CDR loops and the reformantiant antibody. According to the present invention, this problem is removed by the next step which is to identify, by means of a comparison of both of the above-described three-dimensional models of the rodent antibody variable region, any residues from the set of surface exposed amino acid residues of the variable region heavy and light chain pair of the human antibody identified that are within 5 Angstroms of any atom of any residue of the rodent antibody or antibody fragment complementarity deter-

mining regions (CDRs).

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Any residue(s) so identified is then changed back from the human to the original rodent amino acid residue(s).

The results of this method can then be applied to a particular rodent antibody by well known methods. Briefly, genes for the humanized variable heavy and light chain regions are constructed using standard recombinant DNA methods (Sambrook, J., Fritsch, E.F. and Maniatis, T. (1989), Molecular Cloning, Second Edition). For example, a PCR method can be used (Daugherty et al. (1991), Nucleic Acids Research 19, pp. 2471-2476).

Variable heavy chain or variable light chain gene constructs are subcloned into appropriate expression vectors. Suitable expression vectors contain either a human gamma or human kappa constant region gene, a suitable promoter, a sequence coding for a human immunoglobulin leader peptide (for example: met-gly-trp-ser-cys-ile-ile-leu-phe-leu-val-ala-thr-ala-thr (SEQ ID NO:39), Olandi et al. (1989), PNAS **86**, pp. 3833-3837), and a drug selectable marker.

Heavy and light chain expression plasmids can be co-transfected, for example, by electroporation into suitable cells, for example, SP2/0 cells, and selected with an appropriate drug, G418, for example. Screening for intact antibody can be accomplished by ELISA assay. 96-well plates are coated with, for example, goat antihuman kappa chain antibody, and light chains are detected with, for example, goat anti-human antibody conjugated to alkaline phosphatase.

As another approach, light chain constructs are transfected, for example, by electroporation into suitable cells, for example, SP2/0 cells and selected, for example, in hygromycin. Screening for light chain expression can be accomplished by ELISA assay. 96-well plates are coated with, for example, goat anti-human kappa chain antibody, and light chains are detected with, for example, goat anti-human antibody conjugated to alkaline phosphatase.

A light chain producing line is then used as a host to electroporate in the heavy chain construct. The heavy chain plasmid is co-transfected with a plasmid containing the gene coding for another drug marker, for example, neomycin resistance and selected in the presence of the drug G418. Screening for intact antibody is accomplished by ELISA assay. 96-well plates are coated with, for example, goat anti-human Fc and detected with, for example, goat anti-human light chain conjugated to alkaline phosphatase.

#### **EXAMPLE 1 AND COMPARATIVE EXAMPLES**

The superiority of the presently claimed method for determining how to modify a rodent antibody or fragment thereof by resurfacing in order to produce a humanized rodent antibody will now be described by reference to the following example and comparative examples which are illustrative and are not meant to limit the present invention.

#### A) Analysis for Murine Antibodies

In order to determine the positions which are usually accessible on the surface of the  $F_V$  domain of murine antibodies, the accessibility was calculated for twelve Fab x-ray crystallographic structures obtained from the Brookhaven database (Bernstein, F., Koetzle, T., Williams, G., Meyer, E., Brice, M., Rodgers, J., Kennard, O., Shimanouchi, T. and Tasumi, M. (1977), J. Mol. Biol. 112, pp. 535-542). The relative accessibility was calculated using the program MC (Pedersen, J. (1991)), which implements a modified version of the DSSP (Kabsch, W. and Sander, C. (1983), Biopolymers 22, pp. 2257-2637) accessibility calculation routine in which explicit atomic radii are specified for every atom. A residue was defined as being surface accessible when the relative accessibility was greater than 30%. The alignment positions of these residues were conserved in all twelve structures (98% identity). Surface accessible framework positions constitute 40% of the  $F_V$  surface area. The remaining surface accessible residues are in the CDRs and in the interdomain C-terminal region. Figures 3A and 3B show a sequence alignment of the twelve crystal structures, the average relative accessibility, and the 30% accessibility cutoff. Figure 3A shows the alignments relative accessibility for the twelve murine antibody light chains and Figure 3B shows the alignments and relative accessibility for the murine antibody heavy chains.

The surface accessible framework positions were mapped onto a database of unique human and mouse  $F_V$  sequences (see lists at the end of this Example). The frequency of particular residues in each of these positions is shown in Table 1. Only residue frequencies higher than 5% are listed.

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		<u> </u>	Light chain	
5		Position	Human	Mouse
	i	1	D 51 E 34 A 5 S 5	D 76 Q 9 E 6
		3	V 38 Q 24 S 24 Y 6	V 63 Q 22 L 5
		5	T 61 L 37	T 87
		9	P 26 S 26 G 17 A 14 L 7	S 36 A 29 L 17 P 5
		15	P 62 V 25 L 12	L 47 P 30 V 8 A 7
10		18	R 57 S 18 T 13 P 6	R 38 K 22 S 13 Q 12 T 9
		46	P 94	P 82 S 9
		47	G 89	G 71 D 18
		51	K 43 R 31	K 70 Q 13 R 8 T 5
		63	G 91	G 98
		6 <b>6</b>	D 43 S 25 A 9	D 38 A 26 S 26
15	٠. ٠	73	S 96	S 90 I 5
	Note that the state of the stat	76	D 43 T 18 S 16 E 15	D 67 S 15 A 5 K 5
		86	P 44 A 27 S 17 T 8	A 50 P 11 T 8 E 7 Q 6
	. "	87	E 71 D 11 G 7	E 91 D 6
	. * 5	111	K 74 R 12 N 6	K 93
20		115	K 54 L 40	K 87 L 5
20		116	R 60 G 33 S 5	R 89 G 9
		117	Q 50 T 37 E 6 P 6	A 74 Q 14 P 5 R 5
			Heavy chain	
		Position	Human	Mouse
		118	E 47 Q 46	E 59 Q 29 D 10
25		120	Q 83 T 7	Q 68 K 26
		122	V 59 L 15 Q 13	Q 57 V 27 L 5 K 5
	,	126	G 54 A 23 P 18	G 36 P 30 A 29
		127	G 53 E 22 A 14 D 7	E 45 G 43 S 6
		128	L 61 V 31 F 7	L 96
	•	130	K 46 Q 41 E 5	
30		131	P 95	K 52 Q 27 R 17
		131	G 74 S 16 T 7	P 91 A 5
,		136	R 53 K 23 S 17 T 7	G 82 S 17
			G 96	K 66 S 17 R 13
		143		G 98
35		145	T 46 S 32 N 9 I 7	T 63 S 19 N 7 A 5 D 5
		160	P 84 S 10	P 89 H 7
	!	161	G 93	G 71 E 24
		162	K 76 Q 10 R 8	K 50 Q 30 N 10 H 5
		183	D 26 P 25 A 17 Q 10 T 7	E 31 P 22 D 17 A 12 Q 11
		184	S 70 K 9 P 8	K 42 S 37 T 6
40	. – –	186	K 53 Q 22 R 7 N 5	K 83 Q 7
		187	G 66 S 21 T 5	G 62 S 18 D 10
		195	T 30 D 26 N 19 K 7	T 36 K 30 N 26 D 6
		196	S 91	S 76 A 16
		197	K 65 I 8 T 8 R 5	S 46 K 34 Q 11
		208	R 46 T 18 K 17 D 6	T 55 R 26 K 8
45	*	209	A 50 P 21 S 13 T 8	S 67 A 14 T 11
		210	E 46 A 18 D 13 S 9 Z 8 V 5	E 88 D 7
		212	T 91	T 53 S 43

Table 1: Distribution of accessible residues in human VH and VL chain sequences. All of the positions appear to be conserved, which leads to the hyphothesis that immunogeneoity arises from a specific combination of these surface residues. The sequence numbering is explaned in Figures 3A and 3B.

G 17 D 11 P 10 Y 9 V N 8

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None of the entire combinations of surface residues in the human sequences are found in the murine sequences and *vice versa* (see lists at the end of this Example). However, the residues in individual positions appear to be conserved (see Table 1). There are few residues which differ significantly betwe n the species;

these are at positions 54 and 91 of the L chain and 168 and 216 of the H chain. Of these positions only position 216 is a non conservative (V to Y) mutation. Differences between human and murine antigenicities are therefore believed to arise from the combinations of residues in these positions.

In order to determine whether the mouse sequences are more distantly related to human  $F_V$  sequences than to other mouse  $F_V$  sequences, the homology was calculated using a Dayhoff mutation matrix (Dayhoff, M., Barker, W. and Hunt, L. (1983), Meth. Enz. 91, pp. 524-545). The homology was calculated between all the sequences in a pool of both human and mouse sequence patches made up of the surface accessible residues. The data was then represented as a density map (not shown) in which the sequences are plotted against each other. The density map can be used to discriminate "murine surfaces" from "human surfaces".

# B) Reshaping of Antibody N901

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In order to test the resurfacing approach suggested by the above analysis, three humanization experiments were set up. 1) Traditional loop grafting (Verhoeyen, M.E., Saunders, J.A., Broderick, E.L., Eida, S.J. and Badley, R.A. (1991), Disease markers  $\bf 9$ , pp. 3-4) onto a human  $\bf F_V$  framework of known structure (KOL). 2) Resurfacing approach using most similar chain. 3) Resurfacing approach using human sequences with most similar surface residues.

The antibody used was the murine anti-N901 antibody (Griffin et al. (1983), J. Imm. 130, pp. 2947-2951). The anti-N901 antibody (also referred to herein as the "N901 antibody") is available commercially from Coulter Corporation under the name NKH-1.

The alignment of the light chain sequences and heavy chain sequences in Figures 4A and 4B, respectively, show the original N901 antibody and the sequences used in each of the three approaches outlined here.

Figures 4A and 4B show alignments of sequences generated using the three methods of humanization. Sequences are: 1) Original rodent N901. 2+3) KOL (Marquart, M. Deisenhofer, J. and Huber, R. (1980), J. Mol. Biol. 141, pp. 369-391) and reshaped N901 using KOL surface. 4+5) Most homologous sequences, L(KV2F) (Klobeck, H., Meindl, A., Combriato, G., Solomon, A. and Zachau, H. (1985), Nucleic Acids Res., pp. 6499-6513) and H(G36005) (Schroeder, H. and Wang, J. (1990), Proc. Natl. Acad. Sci. U.S.A. 87) and reshaped N901 using these sequences. 6+7) Most homologous with respect to surface residues, L(KV4B) (Klobeck, H., Bronkamp, G., Combriato, G., Mocikat, R., Pohelnz, H. and Zachau, H. (1985), Nucleic Acids Res. 3, pp. 6515-6529) and H(PLO123) (Bird, J., Galili, N., Link, M., Sites, D. and Sklar, J. (1988), J. Exp. Med. 168, pp. 229-245), and reshaped N901 using these sequences. The numbering is the same as used in the antibody modelling program ABM (ABM is a trademark for commercial software, Oxford Molecular Ltd., Oxford, U.K.), which is based on structural conservation and not sequence homology as used by Padlan et al. (Kabat, E.A., Wu, T.T., Reid-Mill r, M., Perry, H.M. and Gottesman, K.S. (1987), Sequences of Proteins of Immunological Interest. U.S. Department of Health and Human Services, Fourth Edition). The sequence changes which have to be introduced in order to reshape N901 with a given sequence are marked with bars, and back-mutations as determined from F<sub>V</sub> models are marked with stars. The sequence homology of a given sequence to N901 is shown in brackets after each sequence.

#### (1) Classical Humanization

In classical humanization the rationale is to graft the rodent CDR's onto a framework of known structure, such that CDR-framework interactions can be accurately monitored by homology modelling. The model of the humanized antibody is compared to that of the original rodent antibody, and possible CDR interacting framework residues are back mutated (marked with '\*' in alignment) in order to retain the three-dimensional shape of the CDR's. In this example the antibody KOL was used, giving a low homology score of only 77 and 46 in the heavy and light chains respectively.

# (2) Most Similar Chain Resurfacing

A database of nonredundant human antibody sequences was compiled from available protein and nucleotide sequences. A total of 164 H and 129 L chains were sampled.

Each of the rodent chains, L and H, were then matched and the most similar human sequence found independently (G36005/KV2F) (Schroeder, H. and Wang, J. (1990), Proc. Natl. Acad. Sci. U.S.A. 87); Klobeck, H., Meindl, A., Combriato, G., Solomon, A. and Zachau, H. (1985), Nucleic Acids Res., pp. 6499-6513). Surface residues, as outlined in Table 1, were then changed in the rodent sequences to match those of the human sequences. Subsequently a model was built of the resurfaced antibody and compared to the model of the original rodent antibody and back mutation of any CDR interacting residues was performed.

# (3) M st Similar Surfac Replac ment Acc rding t th Present Invention

This method is identical to the above method, except that the similarity is calculated only over the surface residues outlined in Table 1 above.

The same procedure of surface mutation and subsequent back mutation was performed as in the previous methods. In this case the chosen sequences were PLO123/KV4B (Bird, J., Galili, N., Link, M., Sites, D. and Sklar, J. (1988), J. Exp. Med. **168**, pp. 229-245); Klobeck, H., Bronkamp, G., Combriato, G., Mocikat, R., Pohelnz, H. and Zachau, H. (1985), Nucleic Acids Res. **3**, pp. 6515-6529).

The following lists show the surface residue patterns in mouse and human light and heavy chain antibody variable regions. The sequences are ordered on similarity to one another. There are no pattern matches between mouse and human sequences although there are matches within a species.

4.

# MOUSE LIGHT CHAIN SURFACE PATCHES

```
:KTSLRPGKGSSDYEKK*
:QTSLRPDKGSSDYEKK*
:QTSLRPDKGSSDHEKK*
:QTSLRPDKGSSDQEKK*
:QSSLRPDKGSSDQEKK*
:QTSLRPDKGSSDPEKK*
:QTSLRPDKGSSDPEKK*
:QTSLRPDKGSSDPEKK*
:QTSLRPDKGSSDPEKK*
:QTSLRPDKGSSDPEKK*
:QTSLRPDKGSSDPEKK*
:QTSLRPDKGSSDPEKK*
              1 KV5E$MOUSE
             2 PL0101
             3 N$1F19L
             4 KV5U$MOUSE
             5 MUSIGLDD
10
             6 PL0220
             7 KV5J$MOUSE
                                                                               (SEQ ID NO: 45)
(SEQ ID NO: 47)
(SEQ ID NO: 48)
(SEQ ID NO: 49)
             8 MUSIGKABB
                                              :QTSLRPDKGSSDPEKT+
             9 MUSIGKCLG
                                     :QTSLRADKGSSDQEKK*
:QTSLRPDKGKSDSEKK*
:QTSLRPARGSSDQEKK*
:QTSLRPGRGSSDPEKK*
:QTSLRPGRGSSDTEKK*
:QTSLRPGKGSSDSEKK*
:QTSLRPGKGSSDSEKK*
:QTSLRPGKGASDADKK*
                                              : QTSLRADKGSSDQEKK*
            10 MUSIGGVJ2
            11 MUSIGKCRN
                                                                                 (SEQ ID NO: 50)
           12 MUSIGKCLP
                                                                                (SEQ ID NO: 51)
           13 MUSIGKACH
                                                                             (SEQ ID NO: 52)
(SEQ ID NO: 53)
(SEQ ID NO: 54)
(SEQ ID NO: 55)
(SEQ ID NO: 56)
(SEQ ID NO: 57)
           14 MUSIGKABE
           15 KV5P$MOUSE
           16 MUSIGRCMK
20
           17 KV3D$MOUSE
                                            : VTALRPGKGASDEDKK+
           18 MUSIGRAAW
                                           : VTALRPGKGASDEEKK*
           19 KV3G$MOUSE
                                            : VTALRPCKGASBABKK+
                                                                                (SEQ ID NO: 58)
           20 KV3E$MOUSE
                                            :VTALRPGKGASDEDDE*
                                                                               (SEQ ID NO: 59)
(SEQ ID NO: 60)
(SEQ ID NO: 61)
(SEQ ID NO: 62)
           21 MUSIGKAAZ
                                            :QTSLRPDKGSSDQETT*
:QMSLTPGKGSSSPEKK*
:VTKVRPGKGDSDSDKK*
           22 MUSIGKONE
           23 MUSIGKBA
           24 KVSASHOUSE
                                            :VTKVRPGKGDSDAEKK*
                                                                                (SEQ ID NO: 63)
           25 MUSIGKV
                                            : VTRVRPGKGDSDAEKK*
                                                                                (SEQ ID NO: 64)
(SEQ ID NO: 65)
(SEQ ID NO: 66)
(SEQ ID NO: 67)
(SEQ ID NO: 68)
           26 HUSIGKCNM
                                            : LTKVRPGKGDSDSEKK*
           27 MUSIGKCLI
                                            :VTKVRPGKGDSDSEQK*
           28 KV5BSMOUSE
                                             :VTKVRPEKGDSDAEKK*
          29 MUSIGKCSA -
                                              :VTXVRPEKGDSDSEKK*
          30 MUSIGKCSR
                                             :VTKVSPGKGDSDAEKK*
                                                                                (SEQ ID NO: 68)
(SEQ ID NO: 69)
(SEQ ID NO: 70)
(SEQ ID NO: 71)
(SEQ ID NO: 72)
(SEQ ID NO: 73)
          31 MUSIGKCST
                                             : VTKVRSGKGESDAEKK+
          32 MUSICKAB
                                            :VTSVKPGKGDSDAEKK*
          33 PL0014
35
                                          : VSSVKPGKGDSDAERT.
          34 MUSIGRACU
                                             *VTSAKPCKCDSDAERK*
          35 PS0023
                                             : VSSAKPGKGDSDAEKK*
                                                                                (SEQ ID NO: 74)
          36 NS2MCPL
                                          :VTSARPGKGDSDAZKK*
                                                                                (SEQ ID NO: 75)
          37 MUSICKADY
                                             :VSPAKPGKGDSDAEKK*
                                                                               (SEQ ID NO: 76)
(SEQ ID NO: 77)
(SEQ ID NO: 78)
(SEQ ID NO: 79)
(SEQ ID NO: 80)
                                          :VTKARPGKGDSDVEKH*
          38 MUSIGKCPP
          39 MUSICIDS
                                             :VTLIPPGKGDSDAEKK*
          40 MUSIGECHS
                                          :VTLLQPGKGDSDAEKK*
          41 B27887
                                         :VTLLQPGKGDSDADKK*
:VTLLQPGKGDSDAERK*
          42 H28840
                                                                                (SEQ ID NO: 81)
          43 KV2G$HOUSE
                                         :VTLLQAGKGDSDAEKK*
:VTLLQPGBGDSDAEKK*
                                                                                (SEQ ID NO: 82)
          44 C27887
                                                                               (SEQ ID NO: 83)
(SEQ ID NO: 84)
(SEQ ID NO: 85)
(SEQ ID NO: 86)
45
          45 JL0029
                                           :LTLLOPGNGDSDAEKK*
          46 Musigkazh
                                           :VTLLQPGKGDSDAEKI*
                                          :VTLPQPGQGDSDPEKK*
:VTLPQPGKGDSDAEKK*
          47 PS0074
          48 MUSIGKCMY
                                                                                (SEQ ID NO: 87)
          49 MUSIGECNX
                                            :VTLPQPGKGDWDAEKK*
                                                                               (SEQ ID NO: 88)
          50 KV2D$MOUSE
50
                                             :VTFLSPGQGDSDAEKK*
                                                                               (SEQ ID NO: 89)
```

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```
51 MUSIGKADW
                                : ESSARPGKGDSDAEKK+
                                                        (SEQ ID NO: 90)
(SEQ ID NO: 91)
      52 KV2A$MOUSE
                                : VTLSSPGQGDSDAEKK*
5
      53 KVIASMOUSE
                                :VTTAKPEKGDSDVEKK*
                                                        (SEQ ID NO:
                                                                     92)
      54 F30534
                                : VTTPKPDKGDSDVEKK*
                                                        (SEQ ID NO: 93)
      55 MUSIGKCLO
                                :VTAPRPGKGASSAEKK*
                                                        (SEQ ID NO: 94)
      56 G27887
                                :VTAPKPGKGTSSAEKK*
                                                        (SEQ ID NO: 95)
      57 MUSIGVKV3
                                : VTTPKPGKGASSAEKK*
                                                        (SEQ ID NO: 96)
      58 MUSIGKCNA
10
                                : VSAPKPGKGASSAEKK*
                                                        (SEQ ID NO: 97)
      59 S03410
                                : VTAPRSGKGASSAEKK*
                                                        (SEQ ID NO: 98)
      60 B32456
                                : VTAPKSGKGASSAEKK*
                                                        (SEQ ID NO: 99)
      61 PL0013
                                :VTAPKPDKGVSSAEKK*
                                                        (SEQ ID NO: 100)
      62 MUSIGLAET
                                :VTAPKSEKGVSSAEKK*
                                                        (SEQ ID NO: 101)
      63 MUSIGVKV1
                                : PTAPKPGKGASSAEKK+
                                                        (SEQ ID NO: 102)
15
     64 KV6K$MOUSE.
                                :LTAPKPGRGVSSAEKK+
                                                        (SEQ ID NO: 103)
     65 G30560
                                : VTAPKSGKGASSAEKR*
                                                        (SEQ ID NO: 104)
     66 MUSIGKBO
                                : VSAPKPGKEGSSAEKK*
                                                        (SEQ ID NO: 105)
     67 MUSIGKCNB
                                : VTAPKPRKGASSAEKK*
                                                        (SEQ ID NO: 106)
      68 H33730
                                :VTFLSPGQGNSDAELP*
                                                        (SEQ ID NO: 107)
      69 MUSIGKCPC
                                :VTFLSPGQGNSDEDLP+
20
                                                        (SEQ ID NO: 108)
      70 KV2C$HOUSE
                                : VTLSSPORGDSDAEKK*
                                                        (SEQ ID NO: 109)
      71 MUSIGLAV
                                : VTAPKSSKGGSSARKK*
                                                        (SEQ ID NO: 110)
      72 MUSIGKCNH
                                : QTSPTPGKGSSDPEKK*
                                                        (SEQ ID NO: 111)
      73 KV5R$MOUSE
                                :QISLIPGKGSYDDEKK*
                                                        (SEQ ID NO: 112)
      74
         KV6ESMOUSE
                                (SEQ ID NO: 113)
25
      75 MUSIGKCNI
                               :VTALKSDKGASSGEKK*
                                                        (SEQ ID NO: 114)
      76 MUSIGLDA
                                : VTPPSPGQGDSAAEKK*
                                                        (SEQ ID NO: 115)
      77 C26317
                                : VTPPSPGQGDSAREKK*
                                                        (SEQ ID NO: 116)
      78 PS0073
                                : VTVRKPGKGDSSDEKK*
                                                        (SEQ ID NO: 117)
      79 A23986
                                : OTSVRLGOGSSDPEKK*
                                                        (SEQ ID NO: 118)
(SEQ ID NO: 119)
      80 MUSIGKABW
                                :KTSLRPWKGSSDSDKK*
30
      81 KV5D$MOUSE
                                : QTDVTQGQGSSQPEKK*
                                                        (SEQ ID NO: 120)
      82 MUSIGE6L
                                :QTAVSQGQGSSQSEXX*
                                                        (SEQ ID NO: 121)
      83 MUSIGECOR
                                :LTAPRTNRGSSDSEKK*
                                                        (SEQ ID NO: 122)
      84 MUSIGKCKN
                                : VTAPSSHRGSSDTEKK*
                                                        (SEQ ID NO: 123)
      85 MUSIGLVD
                                : LLSLSPLKGDSDPKKV*
                                                        (SEQ ID NO: 124)
35
      86 506822
                                : VTAPTPDTGAIKTEKL*
                                                        (SEQ ID NO: 125)
      87 506821
                                :VTIPTPDTGAIKTEKL>
                                                        (SEQ ID NO: 126)
(SEQ ID NO: 127)
     88 MUSIGLAS
                                : AVSPTPDTGAIRTEKL*
     89 MUSIGLAR
                                :AVSPTPDTGAIKTEKL*
                                                        (SEQ ID NO: 128)
     90 LV2B$NOUSE
                                :AVSPTPDTGVIKTEKL*
                                                        (SEQ ID NO: 129)
     91 Musiglan
                                : AVSPTPDTGAIKTEPS*
                                                        (SEQ ID NO: 130)
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# HUMAN LIGHT CHAIN SURFACE PATCHES

```
1 LV4A$HUMAN
                                     :YLPPTPGVIRSTAMKL*
                                                                    (SEQ ID NO: 131)
           2 LV4B$HUMAN
                                       :YLPPTPGVIRSTAMRL*
                                                                    (SEQ ID NO: 132)
(SEQ ID NO: 133)
           3 LV4E$HUMAN
                                        :YLPPTPGLIRSTSMKL*
           4 LV4D$HUMAN
                                        :YLPPTPGLIRSTSVKL*
                                                                    (SEQ ID NO: 134)
           5 LV4C$HUMAN
10
                                       :YLPPTPGVIRSTAEKL*
                                                                    (SEQ ID NO: 135)
           6 LV5A$HUMAN
                                       :YLPPTPGVIRSTAGKL*
                                                                    (SEQ ID NO: 136)
           7 LV7A$HUMAN
                                     : Ylpatpgvvrssagml*
                                                                    (SEQ ID NO: 137)
(SEQ ID NO: 138)
           8 LV2GSHUMAN
                                       :SLPPSPGKVRSTAEKL*
           9 LV2I$HUMAN
                                    : SLPPSPGKVRSTANKL*
                                                                    (SEQ ID NO: 139)
(SEQ ID NO: 140)
          10 N$2RHE
                                      :SLPPRPGKVRSSSEKL*
15
          11 HUMIGLAN
                                        :SLPPRPGKVRSSSDKL*
                                                                    (SEQ ID NO: 141)
          12 LV1ASHUMAN
                                    :SLPPRPGRVRSSEKL*
                                                                    (SEQ ID NO: 142)
          13 LV1B$HUMAN
                                      : Slpprpgkvrssseql*
                                                                    (SEQ ID NO: 143)
          14 LV1FSHUMAN
                                    :SLPPRPGKVRSSSETL*
                                                                    (SEQ ID NO: 144)
          15 LV1C$HUMAN
                                      :SLPPKPGKIRSSTGKL*
                                                                    (SEQ ID NO: 145)
(SEQ ID NO: 146)
                                     :SLPPKPGRIRSSTGKL*
:SLPPKPGKIRSSTGQL*
:SLPPEPGKIRSSTGRL*
         16 A29700
20
          17 HUMIGLAN4
                                                                    (SEQ ID NO: 147)
         18 LV1DSHUMAN
                                                                    (SEQ ID NO: 148)
         19 LV2K$HUMAN
                                        :SLAPSPGKIRSTAPKL*
                                                                    (SEQ ID NO: 149)
         20 LV11$HUMAN
                                       : SLPPRPGKIRSSTGHV*
                                                                    (SEQ ID NO: 150)
(SEQ ID NO: 151)
(SEQ ID NO: 152)
         21 LV2ESHUMAN
                                        : SLRPSPGKVRSTAEKL*
         22 LV2D$HUMAN
25
                                        : SLRPSPGKVRSTADKL*
         23 LV2C$HUNAN
                                        :SLRPSPGKVRSTAENL*
                                                                    (SEQ ID NO: 153)
         24 LV2JSHUMAM
                                       :SLRPSPGKVRSAVEKL+
                                                                    (SEQ ID NO: 154)
                                       : Slpprpgk-rssaekl*
         25 LV1ESHUMAN
                                                                    (SEQ ID NO: 155)
         26 LV2BSHUMAN
                                                                    (SEQ ID NO: 156)
(SEQ ID NO: 157)
(SEQ ID NO: 158)
(SEQ ID NO: 159)
                                       : Slapspckvrstverl*
         27 N$1NCWW
                                      :SLAPSPOKIRSTPOKL*
:SLALSPGKVRSTAEKL*
         28 LV2H$HUMAN
         29 N$3MCG2
                                      : Slplsagkvrstaekl*
: Slapspgkvrstaeyl*
         30 LV2A$HUMAN
                                                                    (SEQ ID NO: 160)
         31 502083
                                      : Slpltpglirstaekl*
                                                                    (SEQ ID NO: 161)
(SEQ ID NO: 162)
(SEQ ID NO: 163)
(SEQ ID NO: 164)
         32 HUMIGLAM2
                                      : SLPLTPRVIRSTARKL*
         33 LV6C$HUMAN
                                      : Flhptpgtdssstekl*
35
         34 LV6DSHUMAN
                                      : Fllptpgtdsssterl*
                                    : Flhptrvtdsssterl*
: Llpptpgtnsssndrl*
         35 LV6ESHUMAN
                                                                    (SEQ ID NO: 165)
         36 LV6B$HUNAN
                                                                    (SEQ ID NO: 166)
         37 HUNIGLESG
                                        :VLPLSPHRIRSESENL*
                                                                    (SEQ ID NO: 167)
(SEQ ID NO: 168)
         38 HUNIGLYC
                                       : SLAPSPAKFRSTAERD*
         39 HUMIGVLLS
                                                                    (SEQ ID NO: 169)
(SEQ ID NO: 170)
(SEQ ID NO: 171)
(SEQ ID NO: 172)
40
                                       :VTAPRPGRIRSDPEKK*
         40 HUNIGKAX
                                     : VTAPRPGRVRSDPEKK+
                                      :VTGPRPGRIRSDPEKK*
:VTGPRPGRIRSDPOKK*
         41 E30609
         42 KV3BSHUMAN
         43 G30607
                                       : VTGPRPGRVRSDPEKK*
                                                                    (SEQ ID NO: 173)
                                                                    (SEQ ID NO: 173)
(SEQ ID NO: 174)
(SEQ ID NO: 175)
(SEQ ID NO: 176)
(SEQ ID NO: 177)
(SEQ ID NO: 178)
         44 KV3MSHUMAN
                                      : VTGPRPGRIRSDPXKK*
45
         45 KV3HSHUMAN
                                      : VTAPRPGRIRSESERK*
         46 KV3KSHUMAN
                                      : VTGPSRGRIRSDPEKK*
        47 KV3P$HUMAM
                                      :VTVPRPSRIRSESERK*
                                      : VTAPGPGRIRSESERK*
         48 B26555
         49 KV1Q$HUMAN
                                       : QTSVRPGRVRSDPERK*
                                                                    (SEQ ID NO: 179)
        50 KV1W$HUMAN
                                       : QTSVRPGKVRSDPERK*
                                                                    (SEQ ID NO: 180)
50
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51 KVIMSHUMAN
                               :QTSVRPGKVRSDPEKK
                                                        (SEQ ID NO: 181)
     52 KV1R$HUMAN
                               :QTSVRPGKVRSEPEKK*
                                                        (SEQ ID NO: 182)
     53
        KV1F$HUMAN
                               :QTSVRPGKVRSEPDKK*
                                                        (SEQ ID NO: 183)
(SEQ ID NO: 184)
5
     54
        KV1G$HUMAN
                               :QTSVRPGKVRAEPEKK*
     55
        KV1K$HUMAN
                               :QTSVRPGKVRSBPZKK*
                                                        (SEQ ID NO: 185)
        KV1D$HUMAN
                               :QTSVRPGKVRSDPBKK*
                                                        (SEQ ID NO: 186)
     57
        KV1H$HUMAN
                               :QTSVRPGQVRSDPERK*
                                                        (SEQ ID NO: 187)
     58 KV1B$HUMAN
                               :QTSVRPGKVRSHPEKK*
                                                        (SEQ ID NO: 188)
10
     59 B27585
                               :QTSVRPGNVRSDPDKK*
                                                        (SEQ ID NO: 189)
     60 NSIREIA
                               :QTSVRPGKVRSDPEKT*
                                                        (SEQ ID NO: 190)
     61 KV1X$HUMAN
                               :QTSVRPGTVRSEPEKK*
                                                        (SEQ ID NO: 191)
     62 KV1L$HUMAN
                               :QTSVRPEKVRSEPDKK*
                                                        (SEQ ID NO: 192)
     63 IMGL38
                               : OTSVRPGKVRSESDKK*
                                                        (SEQ ID NO: 193)
     64
        A27585
                               :QTSVRPGEVRSEPDKK*
                                                        (SEQ ID NO: 194)
15
     :65 KVINSHUMAN
                               :QTSVRPGBVRSBPZRK+
                                                        (SEQ ID NO: 195)
     66 KV1CSHUMAN
                               :QTSVSPGKVRSDPEKK*
                                                        (SEQ ID NO: 196)
        KV1V$HUMAN
     67
                               :QTSVRPGKVNSDPEKK*
                                                        (SEQ ID NO: 197)
        KV1T$HUMAN
     68
                               :QTSVRPGKVRSDPDTK*
                                                        (SEQ ID NO: 198)
        KV1USHUMAN
                               : QTSVRPKKVRSDPZKK*
                                                        (SEQ ID NO: 199)
20
     70 KVIASHUMAN
                               : QTSVRPKKVRFDPEKK*
                                                        (SEQ ID NO: 200)
     71 KV1S$HUMAN
                               :QTSVRSGKVRSEPETK*
                                                        (SEQ ID NO: 201)
     72 KV4A$HUMAN
                               :VTNLRPGKVRSDAEKK+
                                                        (SEQ ID NO: 202)
     73 KV4C$HUMAN
                               : VTDLRPGKVRSDARKK+
                                                        (SEQ ID NO: 203)
        HUMIGK2A1
                               :QTSVSPGNIRSESDKK+
                                                        (SEQ ID NO: 204)
     75
        HUMIGKBA
                               :KTSVTPGKFRSEPEKK*
                                                        (SEQ ID NO: 205)
25
                               :VTLLPPGRVRSDAEKK*
        HUMIGKEC
                                                        (SEQ ID NO: 206)
     77
        KV2B$HUMAN
                               :VTLLPPGEVRSDAEKK*
                                                        (SEQ ID NO: 207)
     78
        KV2D$HUMAN
                               :VTLPPPGZVRSDAERK*
                                                        (SEQ ID NO: 208)
     79
        KV2C$HUMAN
                               :VTLPPPGZVRSBAZNK*
                                                        (SEQ ID NO: 209)
     80 KV2ESHUMAN
                               :VTLPPPQQVRSDAEKK*
                                                        (SEQ ID NO: 210)
30
     81 303876
                               : VTLPPPGQVTSDAEKK*
                                                        (SEQ ID NO: 211)
     82 KV2A$HUMAN
                               : VIILPPAGOVRSDARKR*
                                                        (SEQ ID NO: 212)
     83 HUNIGLAMS
                               : Alspssgqsssaserl*
                                                        (SEQ ID NO: 213)
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# MOUSE HEAVY CHAIN SURFACE PATCHES

```
1 MUSIGHIT
                                : EKVGGLQPGRGTPGKASRGDSQRPES*
                                                                  (SEQ ID NO: 214)
(SEQ ID NO: 215)
       2 MUSIGHIU
                                : EKVGGLQPGRGTPGKVSRGDSQRPES*
       3 MUSIGHIV
10
                                : EKVGGLQPGTGAPGKASRGDSQRPES*
                                                                  (SEQ ID NO: 216)
       4 MUSIGHYM
                                : EKVGGLQPGRGTPGKASKGNSQRAES*
                                                                  (SEQ ID NO: 217)
        PU0003
                                : EKMGGLQPGRGTPGKASKGNSQRAES*
                                                                  (SEQ ID NO: 218)
       6 MUSIGHFO
                                : EKVGGLQPGRGTPGKASKGTSQRAES*
                                                                  (SEQ ID NO: 219)
         A30515
                               : EKVGGLQPGRGTPGKASKGTSQRAET*
                                                                  (SEQ ID NO: 220)
       8
        PL0018
                               : EKVGGLKPGRGTPGKASKGTSQRAET*
                                                                  (SEQ ID NO: 221)
       9 MUSIGHFK
                               : ENVGGLQPGRGTPGKASKGTSQRAET*
15
                                                                  (SEQ ID NO: 222)
     10 MUSIGHPQ
                               : EKVGGLQSGRGTPGKASKGTSQRAET*
                                                                  (SEQ ID NO: 223)
     11 PU0001
                               : EKVGGLQSGRGTPGKASKGTSQRAES*
                                                                  (SEQ ID NO: 224)
     12 E30540
                               : EKVGGLQPGRGTPGKASKGISQRAER+
                                                                  (SEQ ID NO: 225)
     13 HV17$MOUSE
                               : EKVGGLQPGRGTPGKSAKGBSZRAQS+
                                                                  (SEQ ID NO: 226)
        MUSIGHLN
     14
                               : EKVGGLQPGSGTPGKASKGNSQRAES+
                                                                  (SEQ ID NO: 227)
     15
        MUSIGHKG
                               : EKVGGLQPGSGTPGKASKGSSQRAES*
                                                                  (SEQ ID NO: 228)
20
     16 PU0004
                               : EKVGGLQPGRGTPRKASKGNSQRAES+
                                                                  (SEQ ID NO: 229)
        MUSIGHKJ
     17
                               : EKMGNLQPGSGTPGKASKGNSQRPDS+
                                                                  (SEQ ID NO: 230)
     18 HV56$MOUSE
                               : EKVGGLKPGKGTPEKDSKGNARRSET+
                                                                  (SEQ ID NO: 231)
     19
        C27888
                               : ENVGGLKPGKGAPEKDSKGNARRSET+
                                                                  (SEQ ID NO: 232)
     20 MUSIGHAAP
                               : EKVGGLKPGKGTPERDSKGNARRSET*
                                                                  (SEQ ID NO: 233)
     21 PH0097
                               : DKVGGLKPGKGTPEKDSKGNAKRSET+
                                                                  (SEQ ID NO: 234)
     22 E27888
                               : DKVGGLKPGKGTPEKDSKGNAKKSET+
                                                                  (SEQ ID NO: 235)
     23 MUSIGHJB
                               : DKVGGLKPGKGTPDKDNKGNAKKSET*
                                                                  (SEQ ID NO: 236)
     24 MUSIGHADL
                              : EKVGGLTPGKGTPEKDSKGNGRRSET+
                                                                  (SEQ ID NO: 237)
     25 A27888
                               : EMVGGLKPGKGTPEKDSKGNDRRSET*
                                                                  (SEQ ID NO: 238)
     26 H27887
                               : ENVGGLKPGKGTPEKDSKGNDKRSET+
                                                                  (SEQ ID NO:
                                                                              239)
     27
        B27888
                               : EMVGGLKPGKGTPEKDSKGNAKRSET+
                                                                  (SEQ ID NO: 240)
     28 B27889
                               : EQVGGLKPGKGTPEKDSKGHAKKSET+
                                                                  (SEQ ID NO: 241)
     29 D27889
                               : EQVGGLKPGKGTPEKDTKGNAKKSET+
                                                                  (SEQ ID NO: 242)
     30 HV55$MOUSE
                               : EQVGGLKPGKGAPEKDTKGNAKKSET+
                                                                  (SEQ ID NO: 243)
     31 MUSIGHAGT
                               : Exvgglqpgkgtpekdskgnakkset+
                                                                  (SEQ ID NO:
                                                                              244)
     32 MUSIGVHSO
                               : EKVGGLQPGKGTPEKDTKGNAKKSET+
                                                                  (SEQ ID NO:
                                                                              245)
                               : EXVGGLQPGRGTPEXDTKGNAKKSET+
     33 MUSICHIW
                                                                  (SEQ ID NO: 246)
     34 MUSIGHAGE
35
                               : EKVGGLQPGKGSPEKDSKGNAKKSET+
                                                                  (SEQ ID NO: 247)
     35 PH0098
                               : DIDIGGLEPGEGTPEEDSEGNAEQSET+
                                                                  (SEQ
                                                                       ID NO:
                                                                              248)
     36 MUSIGHIM
                               : Equgglapgkgtpdkdskgnakkset+
                                                                  (SEQ ID NO: 249)
     37 MUSIGHAGY
                               : EKVGGLQPGKGTPEKDSKGNAEKSET+
                                                                  (SEQ ID NO: 250)
     38 MUSIGHOET
                               : PQVGDLKPGKGTPEKDTKGNARRSET+
                                                                  (SEQ ID NO:
                                                                              251)
     39 D27888
                               : ENVGDLKPGKGAPEKDSKGNARRSET+
                                                                  (SEQ ID NO: 252)
     40 MUSIGHIP
                               : EQVGGLQPGKGTSDKDSKGNAKKSET+
40
                                                                  (SEQ ID NO: 253)
     41 MUSIGHAGS
                               : EQVGGLQPGKGTPEKDSKGNAKKSGT+
                                                                  (SEQ
                                                                       ID NO:
                                                                              254)
     42 HV16$MOUSE
                               : DQVGGLQPGKGTPEKDTKGNPKRSET+
                                                                  (SEQ ID NO: 255)
     43 B34871
                               : DQVGGLQPGQGTPERNTKGNPKRSDT+
                                                                  (SEQ ID NO: 256)
     44 PH0094
                               : EKVGGLQPGKGTSEKDIKGNAKKSET+
                                                                  (SEQ ID NO: 257)
     45 PH0096
                               : DKVGGLKPGKRTPEKDNKGNAKKSET+
                                                                  (SEQ ID NO: 258)
     46
       MUSIGVH62
                               : DKVGGLKLGKGTPEKDTKGMAKKSFT*
                                                                  (SEQ ID NO: 259)
45
                               : EKVGGLQPGKGTPEKDSKGKANTSET*
     47 MUSIGHAGE
                                                                  (SEQ ID NO: 260)
     48 HV58SMOUSE
                               : EHVGGLKPGKGTPEKDSKGNAGRSET*
                                                                  (SEQ ID NO: 261)
     49 H27888
                               : EOVGGLOPGHGTPEKDTTGNAKRSET+
                                                                  (SEQ ID NO: 262)
     50 HV34SMOUSE
                               : EKEGGLQPGKGTPEKESKGDSKRAET+
                                                                  (SEQ ID NO: 263)
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5
      51 HV33$MOUSE
                                : EKEGGLQPGKGTPEKESKGDSKRPET+
                                                                  (SEQ ID NO: 264)
      52 MUSIGHZAB
                                : EKEGGLQPGKGSPEKESKGDSKRAET+
                                                                  (SEQ ID NO: 265)
      53 NS4FABH
                                : EKDGGLQPGKGTPEKDSKGDSKRVEM+
                                                                  (SEQ ID NO:
                                                                               266)
      54 127888
                                : EQVGGLKPGRGTPEKDTTGDAQRSET*
                                                                  (SEQ ID NO: 267)
      55 G27888
                                : EQVGGLKPGRGTPEKDTTGNAKGSET*
10
                                                                  (SEQ ID NO: 268)
      56 HV59$MOUSE
                                : EKVGGSKPGKGTPEKDSKGNAKTSET*
                                                                  (SEQ ID NO: 269)
      57 MUSIGHOE
                                : SDQGGLKPGKGTPEKDTKGNARRSES+
                                                                  (SEQ ID NO: 270)
      58 N$2FVWH
                                : EKIGGLOPGKGDPGKPSKDNAKRSET*
                                                                  (SEQ ID NO: 271)
      59 MUSIGHJT
                                : EKLGGLQPGKGDPGKPSKDNAKRSET*
                                                                  (SEQ ID NO: 272)
      60 MUSIGHLY
                                : EKLGGLQPGKGDPGKPFKDNAKRSET*
                                                                  (SEQ ID NO: 273)
                                : EKLGGLQPGKGDPGKLMKENAKRSET*
      61 S06816
                                                                  (SEQ ID NO: 274)
15
      62 506817
                                : ENLGGLQPGKGDPGKLKXENAKRPET*
                                                                  (SEQ ID NO: 275)
      63 MUSIGHAAI
                                : EKLGGLOPGNGDLGKPSKDNAKRSET*
                                                                  (SEQ ID NO: 276)
      64 HV42SHOUSE
                                : EKLGPLQLGKGDPGKPSKDDAKRSET*
                                                                   (SEQ ID NO: 277)
      65 MUSICHAAL
                                : EQLGGLQPGGGTPGKPSKDNDKRSET*
                                                                  (SEQ ID NO: 278)
     66 MUSIGHABO
                                : EQLGGLQPGGGTPGKASKDNDKRSET+
                                                                  (SEQ ID NO: 279)
      67
        MUSIGHEG
                                : EQVGGLKARKGTPEKDTTGNAKRSET+
                                                                  (SEQ ID NO: 280)
      68
        MUSIGHWN
20
                                : EMVGVLEPGKGTPEKRQEGNAKRSET*
                                                                   (SEQ ID NO:
                                                                               281)
      69 MUSICKCLT
                                 eqvgglqpkkgspgkdskddsqktet*
                                                                  (SEQ ID NO:
                                                                               282)
      70 MUSIGHZAE
                                : EQVGGLQPKKGSPGKDSKDDSQKTER+
                                                                  (SEQ ID NO:
                                                                               2831
      71 MUSIGHAAD
                                : QQVPELKPGRGTPGKEDKGTSARNDT+
                                                                  (SEQ ID NO:
                                                                               284)
         MUSIGHAAN
                                : QQVPELKPGKGTPGKDDKGTSAKNET+
                                                                   (SEQ ID NO:
                                                                               285)
      73 MUSIGHAMA
                                 QQVPELKPGKGTPGKDDKGTSAKNEN+
                                                                   (SEQ ID NO:
                                                                               286)
      74 MUSIGHXZ
                                : QQKPELKPGKGSPGQEKKGTSSTSET*
                                                                   (SEQ ID NO:
                                                                               287)
25
      75
         A30502
                                : EQQPELKPGKGTPGQEKKGKSSTSES*
                                                                   (SEQ ID NO:
                                                                               288)
      76 MUSIGHAAG
                                : EQQPELRPGKGTPGQEKKGKSSTSES*
                                                                  (SEQ ID NO:
                                                                               2891
      77
         B30502
                                : EQQPELKPGKGTPGQEKKGKSSASES*
                                                                   (SEQ ID NO:
                                                                               290)
      78 MUSIGHADG
                                : EQQPELKPGKGTPGKQKKGKSSTSES*
                                                                   (SEQ ID NO:
                                                                               291)
      79 MUSIGHTV
                                : EQQPELKPGKGTHGKQKKGKSSTSES+
                                                                   (SEQ ID NO:
                                                                               292)
      80 MUSIGHAANA
                                : Eqqpelkpgkgshgkqkkgksstses*
                                                                   (SEQ ID NO: 293)
30
     81 MUSIGHER
                                : EQQPELKPGKGSHGKQKKGKSSASES*
                                                                   (SEQ ID NO:
                                                                               294)
                                : EQQPELKPGKGTHGKQKKGKSSTPES*
     82 MUSIGHAI
                                                                   (SEQ ID NO:
                                                                               295)
                                : Eqopelkpckcthckqkqcksstyes*
     83 MUSIGHALA
                                                                   (SEQ ID NO: 296)
     84 PL0011
                                : EQQPELKPGKGTHGKEKKDKSSTSES*
                                                                   (SEQ ID NO:
                                                                               297)
     85 MUSIGKCLS
                                 BOOAKLKPGKGSHGKQKKGKSSTSES*
                                                                          NO: 2981
                                                                   (SEQ ID
     86 MUSICHADY
                                : EQQPELXPGXGTHGXQXXSNSSTSES+
                                                                   (SEQ ID NO:
                                                                               299)
     87
         MUSIGHWVX
                                : QQQAELRPGKGAPGQEKKGKSSTSES*
                                                                           NO:
                                                                               300)
35
                                                                   (SEQ ID
      88 MUSIGHADO
                                 QQQAELRPGKGAPGQEKKGKSSTSDS+
                                                                   (SEQ ID NO:
                                                                               301)
     89 MUSIGHVBM
                                :QQQAELRPGKGVPGQEKKGKSSTSDS+
                                                                   (SEQ ID NO:
                                                                               3021
     90
        A24672
                                :QQQPELKPGKGAPGKGKKGKSSTSES*
                                                                   (SEQ ID NO:
                                                                               3031
     91 MUSIGHJG
                                : QQQPELRPGKGAPGKGKKDKSSTSES*
                                                                   (SEQ ID NO:
                                                                               304)
     92 JL0044
                                : EQQPEAKPGKGTHGKQKKGKSSTSDS+
                                                                   (SEQ ID NO:
                                                                               3051
     93 MUSIGHBA
                                : QQQAELKPGKGTHGKEKKDKSSTSDS*
                                                                   (SEQ ID NO:
                                                                               306)
40
     94 MUSIGHAGP
                                : QQQAELRPGKGAPGQGKKGKSSTSES*
                                                                   (SEQ ID NO:
                                                                               307)
     95 MUSIGHVBK
                                : QQQAELKPGRGTPGQEKKGKSSTSES*
                                                                           NO:
                                                                               308)
                                                                   (SEQ ID
     96
        A36194
                                : eqqaelragkgtpgqekkgksstses+
                                                                   (SEQ ID NO:
                                                                               3091
     97 MUSIGHVBJ
                                : EQQAELRPGKGTPGQEKKGTSSTSES*
                                                                               310)
                                                                   (SEQ ID NO:
     98 MUSIGHADV
                                :QQQAELRPGKGTPGHEKKGTSSTSES*
                                                                   (SEQ ID NO:
                                                                               311)
     99
         MUSIGHAAT
                                : QQQAELKPGKGTPGHEKKGTSSTSES*
                                                                   (SEQ ID NO: 312)
    100 MUSIGHJL
                                : QQQAELRPGKGTPGHENKGTSSTSES*
                                                                   (SEQ ID NO: 313)
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5 101 MUSIGHABM :QQQAEVRPGKGTPGHEKKGTSSTSES+ (SEQ ID NO: 314) (SEQ ID NO: 315) 102 MUSIGHFU : QQQAELKPGKGTPGHENKGTSSTSES\* 103 MUSIGHZZB :QQQAELRPGKGTPGQQKKGKSSASES\* (SEQ ID NO: 316) 104 HV06SMOUSE : HQQAELKPGKGTPGQQKKGKSSTSES\* (SEQ ID NO: 317) 105 MUSIGHRD : EQQVELRAGKGTPGQEKKGKSSTSES\* 10 (SEQ ID NO: 318) (SEQ ID NO: 319) (SEQ ID NO: 320) 106 MUSIGHVBH : EQQAELRPGKGTPGQEKQGTSSTSES. 107 HV01\$HOUSE : EQQAELRPGKGTPGHDNKGTSSTSES\* 108 MUSIGHADN :QQQAEVRPGKGTPGHEKKGRSSTSES\* 109 HV05SMOUSE (SEQ ID NO: 321) :QQQAELRPGKGTPGQQKKDKSSTSES\* (SEQ ID NO: 322) (SEQ ID NO: 323) 110 MUSIGHAEF :QQQAELKPGKGTPGQQKKDKSSTSES\* 111 MUSIGHAAN :QQQAELKPGKGTPGQQKKDKSSTSDS\* 15 (SEQ ID NO: 324) (SEQ ID NO: 325) (SEQ ID NO: 326) 112 MUSIGHAAB :QQQAELRPGKGSPGQQKKDKSSTSES\* 113 C30560 :QHQAELKPGKGTPGQQKKNKSSTSES+ 114 PS0024 :QQQAELKPGKGTPGQQNKDKSSTSES\* (SEQ ID NO: 327) 115 MUSIGHRG : EQQAELRAGKGIPGQEKKGKSSTSES\* (SEQ ID NO: 328) (SEQ ID NO: 329) 116 MUSIGHAAR : QQQAELKPGKGTPGQEKKSKSSTSES\* "117 MUSIGHLX : QQQSELKPGKGTPGQEKKSKSSTSES\* (SEQ ID NO: 329) (SEQ ID NO: 331) (SEQ ID NO: 332) (SEQ ID NO: 333) (SEQ ID NO: 334) 20 118 HV04\$MOUSE :QQQTELKPGKGTPGQEKKSKSSTSES\* 119 MUSIGHVBG : EQQAELRTGKGTPGQERKGKSSTSES+ 120 MUSIGHMX : QQQAELKPGKGTPGQQKKDKSSTFES+ 121 MUSIGHAAR : EQQAELRPGTGAPGQEKKGKSSTSES\* 122 HV15\$HOUSE : QQQPEVRPGKGTHAKQKKGKSSTSES\* (SEQ ID NO: 335) 123 MUSIGHAAU : QQQPEVRPGKDTHAKQKKGKSSTSES\* (SEQ ID NO: 336) (SEQ ID NO: 337) 124 MUSIGHVBO : QQQAELKPGKGTPEQEKKGRSSTSES\* 25 125 A26405 · : EQQTELRAGKGTPGQEKKGRSSTSEA+ (SEQ ID NO: 338) 126 HV10\$MOUSE :QQQAELKPGKGTPGREKKSKPSTSES\* (SEQ ID NO: 339) (SEQ ID NO: 340) 127 MUSIG3B44 :QQQSELKPGKGTPGREKKSKPSTSES\* 128 MUSIG3B62 : QQRAELKPGKDTPGREKKNXPSTSES+ (SEQ ID NO: 341) 129 HV09\$MOUSE : QQQAELKPGKGTPGREKKSTSSTSES\* (SEQ ID NO: 342) (SEQ ID NO: 343) 130 MUSIGKCLP :QQQAELKPGKGTPGQEKKSTSSTSDS\* 30 131 MUSIGBH : QQQAELRPGKGTPIQQKKDKSSTSES\* (SEQ ID NO: 344) (SEQ ID NO: 345) (SEQ ID NO: 346) 132 HV11\$MOUSE :QQQAEFKPGKGTPGREHRSKPSTSES\* 133 MUSIGHMC :QQQAELRPGKGALGQEKKGKSSTSDS\* 134 MUSIGHAGM : QQQPEVKPGKGAPGKGNTDKSSTSES\* (SEQ ID NO: 347) (SEQ ID NO: 348) (SEQ ID NO: 349) 135 MUSIGHRP : Eqqaevragkgspgqekkgk9st9e9\* 136 MUSIGHVAD : QQLAELKPGKGTPGHEKKGI99T9E9\* 35 137 MUSIGHVAP :QQQAELKPGKGKPEQEKKGTSSTSES\* (SEQ ID NO: 350) (SEQ ID NO: 351) 138 PL0012 : QQQPELRPGKGRHGKENKGKSSTSES\* 139 MUSIGGVD2 :QQQTELRPGRGTTGQERKGKSSTSES\* (SEQ ID NO: 352) 140 506824 : QHQARLKPGKGTPGHENKVTSSTSES\* (SEQ ID NO: 353) (SEQ ID NO: 354) (SEQ ID NO: 355) 141 MUSIGHTS : EQQAELRAGKGTPGQEQKAKSSTSES+ 142 MUSIGHAAR :QQQAELKPGKGTPGQQKTGTSSTTES\* 40 143 MUSIGHHS : QQQABLKPGKGNPGQEKKSTSSASES\*

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144 MUSIGHAXA

146 MUSIGHVBP

148 MUSIGHAYA

150 MUSIGHDZ

149 MUSIGHCP2 -

147 PH0100

145 HV50\$MOUSE

55

: EQQTVLRPGKGTPGQQKKGTSATNES\*

:QQLTELKPGNGTPGQEKKSKSSTSES+

: QQQSVLRPGKGTPGQEKKGTSSTSK9\*

: LOOPVLKPGKGSHGKOKKDKSSTSES+

: EQQPETKPGKGTLGKQKKSKSSTSES\*

: QQQAELKPGQGTPGQEKQQKSSTPEF\*

: EQQAELRPGKGMPEQPKQGTSSTSET+

(SEQ ID NO: 356) (SEQ ID NO: 357)

(SEQ ID NO: 358)

(SEQ ID NO: 359) (SEQ ID NO: 360)

(SEQ ID NO: 361)

(SEQ ID NO: 362) (SEQ ID NO: 363)

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151 MUSIGHNPI
                                  : EQQAELRPGKGNPEQPKQGTSTTSET+
       152 506823
                                                                    (SEQ ID NO: 364)
                                  : EQQAELKPGKGNPEQPKQGTSSTSET*
       153 MUSIGHASA
                                                                    (SEQ ID NO: 365)
                                  : EQQAELKPGKGNPEQPKQDTSSTSET+
       154 S03484
                                                                    (SEQ ID NO:
                                                                                 366)
                                  : EQQAELKPGKGNPEQPKQGTSSTSGT*
       155 MUSIGHVAA
                                                                    (SEQ ID NO:
10
                                                                                 367)
                                  : EQQAEVKPGKGNPEQPKQGTSSTSET+
       156 MUSIGHNPD
                                                                    (SEQ ID NO:
                                                                                 368
                                  : EQQAELRPGKGNPEQPKQVTSSTSET*
       157
          MUSIGHNPB
                                                                    (SEQ ID NO:
                                                                                 369)
                                  : EQQAELRPGKGNPEQPKQITSSTSET+
       158 MUSIGHEC
                                                                    (SEQ ID NO:
                                                                                 370)
                                  : EQQAELRPGRGNPEQPKQVTSSTSET*
      159 MUSIGHNPC
                                                                    (SEQ
                                                                         ID NO:
                                  : EQQAELRPGRGNPEQPKHVTSSTSET+
                                                                                 371)
      160 MUSIGHNPP
                                  : EQQAELRPGKGNTEQPKQVTSSTSET*
                                                                    (SEQ ID NO:
                                                                                 372
      161 MUSICHNPE
                                                                    (SEQ ID NO:
                                  : Eqqaelkpgkgnteqpklitsstset+
                                                                                373)
15
      162 A27635
                                                                    (SEQ
                                                                         ID NO:
                                                                                 374)
                                  :TGQAELRPGKGAPEQGKKGKSSTSDR+
      163 MUSIGHXW
                                                                    (SEQ ID NO:
                                                                                375)
                                  :QYQAELRPGKGTPRQQKKGKSSTSES+
      164 MUSIGHIZA
                                                                    (SEQ
                                                                         ID NO:
                                                                                 376)
                                  : QQQAVLRHGKGTHGQEKKGKSSTSES*
      165 MUSIGHEH
                                                                    (SEQ ID NO:
                                                                                3771
                                 : QQQTKLGPGRGTPGQGRKGKSSTSGS+
                                                                    (SEQ ID NO:
      166 MUSIGHRH
                                                                                378)
                                 : EQQAELRAGKGTPGQEKKGKSSVYPA+
      167 HV00$MOUSE
                                                                    (SEQ ID NO:
                                 : EQQAELKAGKGTPGQQKQGESTRSET+
                                                                                379)
      168 N$1F19H
                                                                    (SEQ ID NO:
20
                                                                                380)
                                 : QQKAELAASKGTPGQEKKGRSSTSES+
      169 MUSIGHZAD
                                                                    (SEQ ID NO:
                                 : QQQTELRPGKGTPGQEKRGKSSNLRL+
                                                                                 381)
      170 B30515
                                                                    (SEQ ID NO:
                                                                                382)
                                 : EKVGGLQGSSFDPGKASKGTSQRAET+
                                                                    (SEQ ID NO:
      171 MUSIGHEB
                                 : EQQADLKLGKGNPEQPKLATPSTSET+
: EQVGGLKPGKGTPDKSDVKDNAKSET+
                                                                                 383)
      172 E27889
                                                                    (SEQ ID NO:
                                                                                384)
      173 MUSIGHAAC
                                                                    (SEQ ID NO:
                                                                                3851
                                 DQQPDLKPSSGSPGHPSKSTSKTTET+
                                                                    (SEQ ID NO:
      174 HV61SMOUSE
                                                                                 386)
                                 : DQQPDLKPSSGSPGNPSKSTSKTTET*
25
      175 MUSIGVHR2
                                                                    (SEQ ID NO:
                                                                                3871
                                 : DQQPDLKPSSGSPGNPSKSTSKTAET+
                                                                    (SEQ ID NO:
      176 PL0100
                                                                                3881
                                 : DQQPGLKPSSGSPGNPSKSTSKTTET+
      177
          MUSIGHAAO
                                                                    (SEQ ID NO:
                                                                                389)
                                 : DQQPGLKPSSGSPGNPSKNTSKTTET+
          MUSIGHGA6
                                                                    (SEQ ID NO:
      178
                                 : DQQPGLKPSSGSPGDPSKTTSKTTET+
                                                                                3901
      179 MUSIGHJY
                                                                    (SEQ ID NO:
                                                                                391)
                                 : DQQPGLKPSSGSPGNPSKTTSKTTET*
      180 MUSIGHGAL
                                                                    (SEQ ID NO: 392)
                                 : DHQPGLKPSSGSPGNPSKNTSKTTET*
                                                                    (SEQ ID NO: 393);
      181 MUSIGHXX
30
                                 : DQQPGLKPSSGSPGMPSRSTSKTTET+
                                                                    (SEQ ID NO:
                                                                                394)
      182 HV62SMOUSE
                                 : Doopglepsagspgnpskstsktaet+
      183 MUSIGHAAGA
                                                                    (SEQ ID NO: 395)
                                 : EQQPGLKPSSGSPGNPSKSTSKTSET*
                                                                    (SEQ ID NO: 396)
      184 MUSIGHGAS
                                 : Doopglkpssgspgnpskntsktiet*
                                                                    (SEQ ID NO: 397)
     185 MUSIGHGA4
                                 : DQQPGLKPSSGSPGDPSKWTSKTPET*
                                                                    (SEQ ID NO:
      186 MUSIGHAGI
                                                                                398)
                                 : EQQPSLKPSSGSPGNPSKSTSKTTET+
                                                                    (SEQ ID NO:
                                                                                399
      187
         PL0102
                                  DQQPGLKPSSGSPGNPSKNTSETTET+
35
                                                                    (SEQ ID NO: 400)
     188 HV46$MOUSE
                                 DOOPGLEPSSGSPGNPSINTSETTZT*
                                                                    (SEQ ID NO: 401)
     189 MUSIGHZT
                                 : EQQPSLKPSSGSPGNPSKSTSKTSET+
                                                                    (SEQ ID NO:
      190 MUSIGHAGD
                                                                                402)
                                 : EQQPSLKPSSGSPGNPSKSTSRTTET+
      191 MUSIGHAGO
                                                                    (SEQ ID NO: 403)
                                 : Doopsladsgspchpskstsktaet+
                                                                    (SEQ ID NO: 404)
     192 MUSIGAM32
                                 : DQQPDLKPSSGPPGNPSKSTSKTTET*
                                                                    (SEQ ID NO: 405)
     193 MUSIGRAFX
                                 : EQQPSLKPSSGSPGKPSKSTSKTNET+
                                                                    (SEQ ID NO: 406)
     194 MUSIGHAGE
40
                                 : EQQPSLKPSSGSPGNPSKSTFKTSET+
                                                                    (SEQ ID NO:
                                                                                407
     195 MUSIGHAGE
                                 : DOOPSLKPSSGSPGNPSKSTSTTSET*
                                                                    (SEQ ID NO:
                                                                                408)
     196 MUSIGHAGE
                                 : EQQLSLKPSSGSPGNPSKSTSKTTET+
                                                                    (SEQ ID NO: 409)
     197 HUSIGHAAN
                                 : QQQPGLKPSPGPPGKPSQSTSKTTET*
                                                                    (SEQ ID NO: 410)
     198 HV43$MOUSE
                                 : QQKPGLAPSGSPGKSTKSNSKQTDT+
                                                                    (SEQ ID NO: 411)
     199 MUSIGMUV1
                                 : QQKPGLAPSSGSPGKSAKSNSKQTDT+
                                                                    (SEQ ID NO: 412)
     200 MUSIGHAZI
                                 : QQKPGLAPSGSPGKSAMSNSKQTDT+
45
                                                                    (SEQ ID NO: 413)
                                 : OOKPGLAPSSGSPGKSAISNSKQTDT*
     201 MUSIGHBP
                                                                    (SEQ ID NO: 414)
     202 MUSIGHZZA
                                 : QQRPGLQPSSGSPGKAA19HSKQSHT*
                                                                    (SEQ ID NO: 415)
     203 MUSICMUV2
                                 : QQKPGLQPSSGSPGKAAISNSKQANT+
                                                                    (SEQ ID NO: 416)
     204 A32456
                                 : QQKPVLAPSSGSPGKSAMSNSKQIDT+
                                                                    (SEQ ID NO: 417)
     205 MUSIGHMB
                                 :QQKPSLQPSSDSPGKAAMSNSKQADT*
                                                                    (SEQ ID NO: 418)
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# HUMAN HEAVY CHAIN SURFACE PATCHES

```
1 HUMIGHVS
                                 : ERVGDLEPGRGIPGKAPKGDSKKIET*
                                                                    (SEQ ID NO: 419)
        2 HUMIGHVR
                                 : ERVGDLEPERGIPGKAPKGDSKKIET*
                                                                    (SEQ ID NO: 420)
        3 H36005
                                 : EQVGGLKPGRGTPGKAPKGDSKKTET*
 10
                                                                    (SEQ ID NO: 421)
          PL0122
                                 : EQVGGLQPGKGTSGKASKGDSKKTET*
                                                                    (SEQ ID NO: 422)
          HV3DSHUMAN
                                 : EQLGGLQPGRGTPGKBSKGDSKRAET*
                                                                    (SEQ ID NO: 423)
        6 HUMIGHAT
                                 : EQLGGLQPGRGTPGKDSKGNSKRAET+
                                                                    (SEQ ID NO: 424)
          B34964
                                 : EQLGGLQPGRGTPGKDSRGNSKRAET+
                                                                    (SEQ
                                                                         ID NO:
                                                                                4251
        8
          A34964
                                 : EQVGGLQPGRGTPGKDSKGNSKRAET+
                                                                    (SEO
                                                                         ID NO: 426)
          PL0123
                                 : EQVGGLQPGRGTPGKDSKGNAKRAET+
                                                                    (SEQ ID NO: 427)
 15
       10 HV3F$HUMAN
                                 : EQVGGLQPGRGTPGKDSKGDSRRAET+
                                                                    (SEQ
                                                                         ID NO:
                                                                                428)
       11
          JL0048
                                 : EQVGGLQPGRGTPGKDSKGHSRRAET+
                                                                    (SEO
                                                                         TD NO:
                                                                                4291
       12 HV3BSHUMAN
                                 : QQVGGLEPGRGTPGKDSKGBSKRAET •
                                                                    (SEQ ID NO: 430)
       13 HUMIGHBV
                                 : EQLGDLQPGRGTPGKASKGNSKRAET+
                                                                    (SEQ
                                                                         ID NO:
                                                                                431)
       14
          HV3ESHUMAN
                                 : EQVGGLQPGRGTTGKDSKGDSKRAET+
                                                                    (SEO
                                                                         ID NO:
                                                                                 432)
       15
         PL0116
                                 : QQVGGVQPGRGTPGKDSKGNSKRAET+
                                                                    (SEQ ID NO:
                                                                                 4331
       16
          HV3K$HUMAN
                                 : QQVGGVQPGRGIPGKDSKGNSKRPET+
 20
                                                                    (SEQ
                                                                         ID NO:
                                                                                434)
          N$2PB4H
                                 : EQVGGVQPGRGIPGKDSKGDSKRPET*
       17
                                                                         ID NO:
                                                                    (SEO
                                                                                435)
       18 HV3I$HUMAN
                                 : QQVGGVQPGRGTPGKDSNGDSKRPET+
                                                                    (SEQ ID NO: 436)
       19
          HV3J$HUMAN
                                 :QKVGGVQPGRGTPGKDSKGNSKRTET*
                                                                    (SEQ
                                                                         ID NO:
                                                                                437)
       20
         HV3G$HUMAN
                                 : QEVGGVZPGRGTPGKBSKGBSKRAET+
                                                                    (SEQ ID NO:
                                                                                438)
          HV3H$HUMAN
       21
                                 : EQLGGLQPGRGTPGKDSNGDSKOAZT+
                                                                    (SEQ
                                                                         ID NO: 439)
      22 HV3O$HUMAN
                                 : EQLGGLOPGRGSPGKDTNGDSKEAZT*
 25
                                                                    (SEQ
                                                                         ID NO:
                                                                                440)
                                 : AQLGGLQPGRGTPGKDSNGDSKQAZS*
       23
         HV3N$HUMAN
                                                                    (SEQ ID NO: 441)
       24 HV3RSHUMAN
                                 : EQLGGLQPGRGTPGKVSQGDSKQAZT*
                                                                    (SEQ
                                                                         ID NO: 442)
       25 HV3P$HUMAN
                                 : EQVGGLQPGRGTPGKVSQGDSKEPZT+
                                                                    (SEQ
                                                                         ID NO:
                                                                                443)
       26
         HUNIGHCV
                                 : EQLGGLQPERGTPGKESKGNSNRAET+
                                                                    (SEQ ID NO: 444)
       27 HV3TSHUMAN
                                 : EQVGDLQPGRGBPGKDSKGNAKRVET+
                                                                    (SEQ
                                                                         ID NO:
                                                                                445)
      28 HVJU$HUMAN
                                 : EQVGDLQPGRGNPGKDSKGNAQRPET+
                                                                    (SEQ
                                                                         ID NO:
                                                                                 446)
 30
      29 PL0098
                                 : QQVGGVQPGRGTLGKDSKGNSKRAET+
                                                                         ID NO:
                                                                    (SEQ
                                                                                 447)
      30 HV3H$HUHAN
                                 :QZVGGAZPGRGSPGKASKGBSKRAET+
                                                                    (SEQ
                                                                         ID NO:
                                                                                448)
      31 HV3A$HUMAM
                                 : QQVGGLKPGRGSPGKDSKGNAQRTZT*
                                                                         ID NO:
                                                                    (SEQ
                                                                                4491
                                 : DQVGGLKPGRGTPGKHSNGDSKTP2T*
      32
         HV3S$HUNAN
                                                                    (SEQ
                                                                         ID NO:
                                                                                450)
      33 HUMIGHAN
                                 : EQLGGLQPGRGTSREDSKGMSKRAET*
                                                                    (SEQ
                                                                         ID NO:
                                                                                451)
      34 HV3Q$HUHAM
                                 : DOVGALOPGRGTPGKDSQADSKEAZT*
                                                                    (SEQ
                                                                         ID NO: 452)
 35
      35
         A36040
                                 : EQLGGLOPGRGTPGK----VEGSVET*
                                                                    (SEQ ID NO: 453)
      36 HUNIGHAM
                                 : EQVGAFQPGRGNSGKASKGDSKRPDT+
                                                                    (SEQ
                                                                         ID NO: 454)
         HUMIGHAO
      37
                                 : EQVGAPQPGKGNSGKASKGDSKRPDT+
                                                                    (SEQ ID NO: 455)
      38
         HUNIGHAR
                                 : EQVGAFQPGKGNSGKASKGDSNRPDT*
                                                                    (SEQ ID NO: 456)
      39
         HVOLSHUMAN
                                 : QQVGGVQAGRANPGKDSRGISKRTET*
                                                                    (SEQ
                                                                         ID NO: 457)
      40 HV1A$HUMAN
                                 : QQVAEVKPGKGTPGQQKQGESTRSET+
                                                                    (SEQ ID NO: 458)
                                 : QQVAEVKPGKGTPGQQKQGTSTRSET+
      41
         A32483
                                                                    (SEQ ID NO: 459)
 40
                                 : QQVAEVKPGKGTPGQQKQGTSARSET*
      42 HUMIGHAY
                                                                    (SEQ ID NO: 460)
      43 HUMIGHCU
                                 : QQVAEVKPGKGTPGQQKQGTSIRSDT+
                                                                    (SEQ ID NO: 461)
         HUMIGHBS
                                 : QQVAEVEPGKGTPGQEKQGTSIRSDT*
      44
                                                                    (SEQ ID NO: 462)
      45 HUMIGVHLS
                                 : QQVAEVKPGKGTPGQQMQGTSTRSDT+
                                                                    (SEQ ID NO: 463)
                                 : QQVGEVEPGRGTPGQQKQDTSTRSDT+
                                                                    (SEQ ID NO: 464)
      46
         HUNIGHBX
      47
         HV1C$HUMAN
                                 : QQVAEVKPGRGTPGHPRQGASFRSDS*
                                                                    (SEQ ID NO: 465)
45
         H34964
                                 : QQVSELKPGKGTPGQQGTGTSVKAET*
      48
                                                                    (SEQ
                                                                         ID NO: 466)
         HUMIGHCY
                                 * Eqvaevkpgkgspgkpsqgks ikast*
      49
                                                                    (SEQ ID NO: 467)
         PL0119
                                 : Eqvaevkpgrgspgkpsqgksikast*
                                                                    (SEQ ID NO: 468)
      50
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	51 HV1FSHUMAN	:QQVAEVKPGRGDPGRPRQASSTISAT+	(SEQ ID NO: 46	59)
5	52 D34964	: EQVAEVPOGKGRPGKSLOGKSLKAST+	•	70).
		: OOMAEVKPGRGTPGKPGVVPSFFSET*		71)
	53 HV1D\$HUHAN		, –	•
	54 HVIESHUMAN	:QQVAEVKPGRGTPGRYIWEPSFFNEG*		72)
	55 JL0047	: QQQAGLKPSSGSPGKPSKSTSKTAAT*	, · · · ·	73)
	56 HUMIGHBW	: QQQPGLKPSSGSPGKPSKSTSKTAAT*	(SEQ ID NO: 47	74)
	57 E34964	: QQQPGLKPSSGSPGKPSKSTSNTAAT*	(SEQ ID NO: 47	75)
10	58 HUMIGHCW	:QQQPGLKPSSGSAGKPSKSTSKTAAT*	(SEQ ID NO: 47	76)
	59 HV2FSHUHAN	: RQQPGLKPSSGPPGKPSRGTSRSAAT*	(SEQ ID NO: 47	77)
	60 HV2ISHUMAN	: QQQAGLKPSSGSPGRTSKSTSKTAAT*	(SEQ ID NO: 47	78)
	61 HV2GSHUMAN	: QQEPGLRPSSGTPGRTPRSTSKTAAT*	(SEO ID NO: 47	79)
	62 NS3FABH	: XQEPGLRPSSGSPGRTPRSTSKTAAT*	(SEQ ID NO: 48	80)
	63 PS0091	: QQQPGLKPSSGSPSRVSKSTSKTPET*	·	81)
15	64 HUMIGHDA	: QHQAGLKRSSGPPGKPSTSTSKTAAT*	` = = = -	82)
13	65 A26555	: Zoesglkptsgspgkpsksrskaada*		83)
	66 HV2ESHUMAN	:OTKPTLKPTTGSPGRPSKSTSKDPVT*	,	84)
	67 HV2DSHUMAN	:OTKPTLKPTTGSPGKPSRSTSRDPVS*		85)
	68 A36005	: ETRPALKPTTGSPGKTSKTTSKDPVT*	/	86)
		: ONRPALKATTGSPGKTSETTSKDPAT*		87)
		:QTTPALKPKTGSPGKTSRTDSKNPVT*	,	•
20	70 HV2ASHUMAN		, · · · · ·	88)
	71 HV2C\$HUHAN	:QTRPALRPTTGSPGEASETTSKGPGT*	,	89)
	72 HV2B\$HUMAN	: QTRPALKPITGSPGKTSETTSRDTAY*	,	90)
	73 JL0049	: Legyqlwggrgisrkyakgngkrdes*	(SEQ ID NO: 4	91)

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#### **EXAMPLE 2**

# DETAILED DESCRIPTION OF METHOD FOR CONSTRUCTING THREE-DIMENSIONAL MODEL OF ANTIBODY VARIABLE REGION

The references cited in the text below are listed at the end of this Example.

The first antibody Fab structure was determined in 1972. Since then, no more than about twelve Fab structures have been published, a number that represents a very small fraction of the total antibody repertoire (>108 antibodies). To understand the molecular basis of this antibody diversity will require knowledge of either a large number of x-ray structures, or the rules by which combining site topography is governed. The development of such prediction rules has now reached the point where variable regions of antibodies can be modelled to an accuracy approaching that of the medium resolution x-ray structure.

The interaction of an antibody with its cognate antigen is one of the most widely accepted paradigms of molecular recognition. To understand the antibody-antigen interaction in atomic detail requires knowledge of the three-dimensional structure of antibodies and of their antigen complexes. Traditionally such information has come from x-ray crystallographic studies (see Davies et al. for review (Davies et al., 1988)).

The modelling of antibody combining sites was first attempted by Padlan & Davies (Padlan et al., 1976) at a time when very few antibody structures were known. Nonetheless, Padlan and colleagues recognized that the key lay in high structural homology that existed within the  $\beta$ -sheet framework regions of different antibody variable domains. The antigen combining site is formed by the juxtaposition of six interstrand loops, or CDRs (Complementarity Determining Regions) (Kabat et al., 1987), on this framework. If the framework could be modelled by homology then it might be possible to model the CDRs in the same way. Padlan and Davies (Padlan et al., 1976) reasoned that CDR length was the important determinant of backbone conformation though the number of antibody structures was insufficient to thoroughly test this maximum overlap procedure (MOP). This notion was not picked up again until the early 1980's when Pedersen and Rees proposed a similar approach to modelling antibody combining sites based on a more extensive analysis of antibody structures (de la Pas et al., 1986).

Those ess ntially knowledge-based procedures are best exemplified for antibodies by the work of Chothia & Lesk (Chothia et al., 1986) who, in 1986, extended and modified the MOP procedur—by introducing the concept of "key" residues. These residues allow the further subdivision of CDRs of the same length into "canonical" structures which differ in having residues at specified positions that, through packing, hydrogen bonding or the ability to assume unusual values of the torsion angels  $\phi$ ,  $\psi$  and  $\omega$ , determine the precise CDR conformation

(Chothia et al., 1989). Similar knowledge-based methods have been proposed for predicting loop conformations in general (Thornton et al., 1988; Tramontano et al., 1989). These methods rely on the crystallographic database of protein structures. However, none of the above knowledge-based methods has been totally successful. In particular, the MOP or canonical structure approaches have succeeded in modelling only five of the six CDRs. This stems from the fact that the third CDR of the heavy chain, H3, is more variable in sequence, length and structure than any of the other CDRs.

To deal with this problem several groups have attempted to use ab initio methods to model the combining site (Bruccoleri and Karplus, 1987). The requirement with such methods is that the total allowable conformational space accessible to a particular CDR is sampled. Typical of purely geometric approaches is that of Go & Sheraga (Go and Sheraga, 1970) and more recently Palmer & Sheraga (Palmer and Sheraga, 1991), where the problem is reduced to one in which the central region of the polypeptide backbone, having characteristic bond length and bond angles, is constructed between the end points of the loop (CDR if an antibody loop) by a "chain closure" algorithm. In a modification of this algorithm, Bruccoleri & Karplus (Bruccoleri and Karplus, 1987) introduced an energy minimization procedure which greatly expanded the domain of conformational space searched during the chain closure procedure. This modification is incorporated into the conformational search program CONGEN (Bruccoleri and Karplus, 1987), which also allows the user to choose any set of standard bond length and bond angels such as the CHARMM (Brooks et al., 1983) standard geometry parameter sets. Other approaches such as minimization (Moult and James, 1986), or molecular dynamics (Fine et al., 1986) either fail to saturate conformational space or are unable to deal with the problem of long CDRs. Whichever of the ab initio methods is employed however, the problem is one of defining the selection criteria in such a way as to allow the unambiguous identification of the correct structure (in this context correct is defined by reference to an appropriate X-ray structure) within the ensemble of candidates, for every CDR. To date this has not been possible.

Recently a more holistic approach has been taken to the modelling of CDRs which combines the advantages of knowledge-based and *ab initio* methods in a single algorithm known as CAMAL (Combined Algorithm for Modelling Antibody Loops) (Martin et al., 1989; Martin et al., 1991). Previously this algorithm has been used to model individual CDRs in the presence of the crystal structure conformations of the other five. As is demonstrated below, CAMAL is able to predict the backbone conformations of all six CDRs of the antibody combining site to an accuracy approaching that of medium resolution x-ray structures. In addition the algorithm includes a procedure for selecting and fitting together the light and heavy chain framework regions prior to generation of CDR conformations, thus making possible the prediction of the entire variable region. Furthermore a new Monte Carlo (MC) simulated annealing method has been developed for the determination of sidechain conformations.

# The Framework Region

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Antibody framework regions consist of conserved  $\beta$ -strands that form the  $\beta$ -barrel structure characteristic of immunoglobulin V-type regions. In the procedure described here each V-region is built from a database of known antibody structures, using sequence homology for selection of the light (L) and heavy (H) chain V-domains. The two domains are then paired by least squares fitting on the most conserved strands of the antibody  $\beta$ -barrel (Table 2 and Figures 5 & 6. The strand orientations were determined by analyzing the barrels of known antibody crystal structures. Eight antibodies were analyzed using a multiple structure fitting program as follows. Seven structures were fitted onto one of the set selected at random and mean coordinates were calculated. All eight structures were then fitted onto these mean coordinates and new mean coordinates determined. This procedure was iterated until the mean coordinate set converged (5-10 cycles). The variance for the mean coordinates at each barrel point (N,C $\alpha$ ,C) was calculated. In Figure 5 this variance is plotted against the projected positions of these points onto the conjugate axis of the barrel.

Strand 8 and all but two residues of strand 7 in both light and heavy chains were eliminated as they showed deviations greater than 3 $\sigma$  (standard deviation units) from the mean coordinates. These two strands comprised the takeoff points of CDR H3, and suggests that any knowledge-based prediction of CDR H3 would have to account not only for sequence and length variation in the CDR itself, but also for the position of the participating strands. The remaining mean coordinates were used as a scaffold onto which the L and H chains were fitted. Strands 7 and 8 in the final framework were obtained from the database structure used in the construction. The framework strands are marked + in the multialignment in Table 2.

The sidechains were then replaced using a 'maximum overlap' method, in which sidechain templates were fitted on backbone atoms with the sidechain torsion angles being adjusted to match those of equivalent torsions in the par int sidechain.

# Th C mbining Site

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The procedure for predicting the structure of combining sites combines a database search with a conformational search procedure. The architecture of the program suite to perform this task is outlined in Figure 7.

The database search utilizes distance constraints for each of the six CDR loops determined from known antibody structures. These constraints were determined by calculating  $C\alpha$ - $C\alpha$  distances within known loops and using a search range of  $\bar{x} + 3.5\sigma$  (the mean  $\pm$  3.5 standard deviation units). A database containing all the proteins in the Brookhaven Protein Databank (Bernstein et al., 1977) is then searched for fragments which satisfy the constraints for a loop of the required length. The middle section of the loop is then deleted and reconstructed using the conformational search program CONGEN (Bruccoleri and Karplus, 1987). For loops of six or seven residues, the structure database appears to saturate the conformational space available to the backbone adequately and only sidechains are built by conformational search. Loops shorter than six residues are built by conformational search alone since this is computationally feasible and the number of loops selected from the database becomes unacceptably large as loop length decreases.

When modelling a complete combining site, loops of 6 or more residues are modelled individually with the other loops absent. If the loops are built consecutively, small errors can accumulate leading to a poor result (Martin, 1990). All the loop conformations are then evaluated using a solvent modified potential, which excludes the attractive van der Waals and electrostatic terms of the non-bonded energy function contained within the GROMOS (Åqvist et al., 1985) potential. The lowest five energy conformations are selected and filtered using a "structurally determining residue" algorithm (FILTER), based on backbone torsion angles observed in the original database loops. Since the database search is not used for the shortest loops of 5 residues or fewer, the FILTER algorithm cannot be used. Energy is thus the only available selection criterion and the short loops are built last, in the presence of the longer loops.

#### Side Chains

The determination of sidechain positions was previously done using the iterative sidechain determination algorithm described by Bruccoleri et al. (Bruccoleri and Karplus, 1987). Unfortunately the CHARMM (Brooks et al., 1983) force field fails to select the correct conformations of exposed hydrophobic sidechains. There is no penalty for having an exposed uncharged atom, without solvent present. CONGEN is also unable to saturate the conformational space for a large number of sidechains (more than 6 residues).

Recently Lee et al. (Lee and Levitt, 1991; Lee and Subbiah, 1991) has proposed a method for searching conformational space for a large number of sidechains using MC simulated annealing. A simple energy function is used for the evaluation of conformations generated by a biased random walk:

$$E = \sum_{i=1}^{n} \epsilon_{o} ((\frac{r_{o}}{r})^{6} - 2(\frac{r_{o}}{r})^{12}) + \kappa_{o} \cdot COS(3\omega)$$

Where the first term is a simple *Lennard-Jones* potential which evaluates the non-bonded contacts between the atoms in a given molecule, the second term is a simple torsional term which only applies to C-C bonds. The torsional term biases the function towards  $60^{\circ}$  rotamers.  $\epsilon_{o}$  and  $\kappa_{o}$  are constants. The metropolis function:

$$P = C^{\frac{-\delta E}{T}}$$

is used to evaluate the energy function. Any move which results in a decrease in energy is accepted, and any move which results in a positive  $\delta E$  is only accepted with the probability P. This simple method can be used to search the large conformational space defined by a set of torsion angles in amino-acid sidechains, and find or define the global minimum which exist for a set of sidechains. T is the simulation temperature.

When searching sidechain conformations using this method the simulation system usually gets trapped in an energetic minima well before the global minimum is encountered, at a high temperature, without the solution space having been searched sufficiently. This problem can be solved by truncating the *Lennard-Jones* potential, thus allowing atoms to pass through each other. In reality this function would converge towards infinity when the distance r between the atoms approach s zero.

The evaluation of sidechain conformations generat d is done sol ly on the basis of energy, for internal (core) residues, sinc good van d r Waal's interactions are considered to be qual to a good packing of the sidechains. The situation becomes more complicated when trying to predict the conformation of surface residues. The lowest van der Waal's interaction is obtained by a combination of sidechain conformations which minimize the overlap of atoms, this means that the lowest nergy is obtained with extended conformations of

sidechains, without considering good packing of sidechains.

Using the fact that hydrophobic, bulky residues will be shielded by the hydrophilic sidechains, and will be buried in the surface, it is possible to generate a simple function which will evaluate these macroscopic observations. These functions can either be implemented in the objective evaluation function of the Monte Carlo simulation, or as is done here, added as a post processing step. Including an accessibility/hydrophobicity term in the evaluation function would slow down the calculation considerably, hence the term has been added as a post processing function. The function used is a sum of the product of relative exposed surface area multiplied by the residual hydrophobicities. The hydrophobicities used are taken from Cornette et al. (Cornette et al., 1987).

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$$f_{conformation} = \sum_{i=1}^{n} -A_{irel} \cdot H_{irel}$$

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n is the number of sidechains reconstructed. The surface area is calculated using the tesselated icosahedron approach (Chau and Dean, 1987), which is not very precise (0.1 percent), but is able to evaluate a large number of conformations. The function is evaluated for the final 2,000 conformations and the lowest value conformation selected as the best.

Using this simple approach it is possible to integrate over a large phase space with many degrees of freedom, and get a complete sampling of the space.

# Predicted Structures of an Anti-hapten, Anti-peptide and Two Anti-protein Antibodies

In the following section the predicted structures of four different antibody  $F_V$  regions are presented and analyzed. The antibodies are:

- Gloop-2 (Darsley and Rees, 1985), an anti-lysozyme antibody whose Fab structure was determined by Jeffrey et al., (Jeffrey et al., 1991) and which was used as a learning exercise during the developm into of CAMAL.
- D1.3 (Amit et al., 1986), an anti-lysozyme antibody whose uncomplexed F<sub>V</sub> coordinates were supplied by R. Poljak et al. after the model coordinates had been deposited.
- 36-71 (Rose et al., 1990), an anti-phenylarsonate antibody whose Fab structure was carried out by D.
   R. Rose, et al., and whose coordinates were obtained after the model coordinates had been deposit d.
- 3D6 (Grunow et al., 1988), an anti-protein (GP41 of HIV) antibody whose Fab structure was carried out by D. Carter et al. (Carter, 1991) and whose coordinates were obtained after the model coordinates had been deposited. For this antibody, the model was generated using the canonical loop method of Chothia & Lesk (Chothia et al., 1989; Chothia et al., 1986) for CDRs L1, L2, H1 and H2, while L3 and H3, which cannot be modelled using canonical structures, were constructed using CAMAL.

All four models were subjected to both restrained and unrestrained energy minimization using the DIS-COVER (TM Biosym Technology) potential with 300 cycles of steepest descents, followed by conjugate gradient minimization until convergence to within 0.01 Kcal occurred.

The resolution and R-factors of the x-ray structures are given in Table 3 together with the parent frameworks selected in building the models. The structures and models were compared by global fits of the loops. The  $\beta$ -barrel strands 1 to 6, as described above, were least squares fitted and the RMS deviation was then calculated over the loops. The backbone (N,C $\alpha$ ,C) RMS values for fitting model and crystal structure frameworks were between 0.4 and 0.9 Å, illustrating the conservation of the core  $\beta$ -barrel. Using all eight strands RMS deviations between 0.6 and 1.2 Å were observed.

Global fits (Table 4) give a more realistic measure of the accuracy of the model than a local least-squares fit over the loops since they account for the overall positioning of the loops in the context of the  $F_V$  structure. Local fits, which give lower RMS deviations, are also shown in Table 4. Differences between local and global RMS deviations arise from differences in  $V_H/V_L$  domain packing and differences in loop 'take off' angles and positions.

Table 5 shows the canonical loops selected from modelling 3D6. Backbone structures of the modelled CDRs, superimposed on the x-ray structures after global fitting are shown in Figure 8. General features and points of interest for each of the six CDRs ar discussed below.

# **Analysis of the CDR Regions**

During the comparison of CDR conformations in the V-region models and the x-ray Fab structures it was observed that at certain positions in a CDR, the peptide backbone may adopt—ither of two conformations by undergoing a "peptide flip" (1,4 shift). This phenomenon is also seen in type 2  $\beta$ -turns (Paul et al., 1990). Dynamics simulations of  $\beta$ -turns show that the transformation energy between  $\phi 1$  = -00,  $\psi 1$  = -30,  $\phi 2$  = -90,  $\psi 2$  = 0 and  $\phi 1$  = -00,  $\psi 1$  = 120,  $\phi 2$  = 90,  $\psi 2$  = 0 has a maximum value of 5 kcal (Paul et al., 1990). This is low enough to allow selection of either conformation. The peptide flip is observed within several canonical classes (as described by Chothia et al. (Chothia et al., 1989)) and the hydrogen bonding pattern used to determine the conformation of a canonical class does not disallow the peptide flip. Any modelling procedure should therefore take these, or any other multiple conformations, into consideration where the transformation energies ar sufficiently low to permit population of the different conformational forms. Table 6 shows an example of the "peptide-flip" phenomenon from the crystallographic database of antibody structures. It should be noted that a single crystal structure will not show multiple conformations since the crystallization will 'freeze out' one of the conformations. During the modelling procedure the two populations of conformers are easily extracted from a set of *ab initio* generated loops, by using a torsional clustering algorithm.

#### CDR-L1

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In Gloop-2 and D1.3, all five low energy conformations were very similar with RMS deviations differing by less than 0.25 Å (backbone) and 0.35 Å (all atoms). The FILTER algorithm was unable to distinguish between the conformations and the lowest energy structure was selected.

Although CDR-L1 of 3D6 was originally built using the canonical loop from HyHEL-10, the mid-section was rebuilt by conformational search, for the following reason. HyHEL-10 and REI CDR-L1 loops are placed in the same canonical ensemble (Chothia et al., 1989) although they contain a 1-4 shift (peptide flip) relative to one another between the fifth and eighth residues of the loop (residues 28-31) (see Table 6).

36-71 shows the same 1-4 shift between the model and crystal structure CDRs. Both crystal structure and model were compared with other loops of the same canonical class as defined by Chothia et al. (Chothia et al., 1989). It was found that the hydrogen bonding pattern which determines the conformation was conserved.

#### CDR-L2

CDR-L2 of D1.3 has two adjacent threonines (49, 50) which in the x-ray structure are packed against the tyrosine at the fourth position of CDR-H3, thus minimizing the exposed hydrophobic sidechains. In the unminimized model the threonine sidechains are exposed to the solvent, but after energy minimization, this packing is observed.

#### CDR-L3

In Gloop-2, D1.3 and 36-71 the proline at the seventh position in the loop is correctly predicted in the *cis* conformation. It has previously been suggested that the conformation of CDR-L3 is dictated by the presence of a proline in position 8 or 9 (Chothia et al., 1989) within the loop. 3D6 does not have a proline in either position. Only 7 out of 290 CDR-L3 sequences (Kabat et al., 1987) lack a proline at both positions and in all of the published x-ray structures this proline is present. This is an example of a situation where either a new canonical class may need to be defined or where the canonical rule breaks down altogether, and an alternative method must be employed.

The 3D6 L3 loop is 7 residues in length and was built using database loops alone where conformational space is saturated by means of fragments selected from the crystallographic database (Global RMS: 2.01 Å, N,C $\alpha$ ,C), and by using CAMAL (Construction: Q[Q(YNS)Y]S, Global RMS: 1.97 Å, N,C $\alpha$ ,C). The similarity of the structures generated by the two procedures illustrates the utility of the database search and suggests that, for shorter loops it is capable of saturating the available conformational space.

#### CDR-H1

Using the Kabat and Wu definition of CDR-H1 places this loop as an extension of the β-sheet. The extended nature of this stretch of peptide limits its conformational flexibility and CDR-H1 is gen rally modelled accurately (Martin et al., 1989; Chothia et al., 1989).

In Gloop-2 and D1.3, the Phe or Tyr sidechain at the second position in the loop is poorly placed and packs against Leu at the penultimate position in HFR1 (see Table 2). 36-71 has a well-placed Asn at this position, rather than the more common bulky hydrophobic sidechain.

#### CDR-H2

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CDR-H2 of 36-71 is similar in sequence to F19.9 (Strong et al., 1991), (36-71: YNNPGNGYIA (SEQ ID NO:492); F19.9: YINPGKGYLS (SEQ ID NO:493)). While the structurally determining residues specified by Chothia and Lesk (Chothia et al., 1989) are conserved, the backbone conformations are different: F19.9 has a bulge at the -PGN- Gly, compared with 36-71, giving the loop a 'kink' in the middle. The model of 36-71 shows a 1-4 shift, though the sidechains are still well placed.

In Gloop-2, the all atom RMS deviation is poor (3.00 Å) (Jeffrey et al., 1991) when compared with the P2<sub>1</sub> crystal structure, owing to rotations of the Phe at position 3 in the loop and Tyr at position 10 by approximately 120° about the  $\chi_2$  torsion angle. Gloop-2 has been solved in two different crystal forms, P2<sub>1</sub> and P1 (Jeffrey et al., 1991; Jeffrey, 1989). When compared with the P1 structure, the sidechains are placed almost perfectly and the all atom RMS (global fit) drops to 2.23 Å.

This concerted sidechain motion between crystal forms illustrates the effects of crystallization conditions on surface sidechain placement. Even though surface sidechains may show low temperature factors indicating low mobility in the crystal, their mobility in solution may be high. In the Gloop-2 P1 structure, the mean sidechain temperature factor for the  $F_V$  domain is 13.46 ( $\sigma$  = 8.20) while the sidechains of these two residues of H2 show mean temperature factors of 5.56 ( $\sigma$  = 0.68) for the Phe at position 3 and 7.10 ( $\sigma$  = 1.73) for the Tyr at position 10.

# CDR-H3

CDR-H3 is the most variable of the six CDR's with all lengths up to 21 residues being represented in Kabat et al., (Kabat et al., 1987). This extreme variability results from V-D-J splicing (Schilling et al., 1980) and has always been a problem when attempting to model antibodies. Such loops may be divided into short (up to 7 residues), medium (up to 14 residues) and long (15 or more residues). Using the CAMAL procedure, short and medium CDR-H3's can be modelled as accurately as other CDR's of similar lengths. Although long CDR-H3's are more difficult and cannot, at present, be built to the same accuracy, the chain trace is still correct.

It is unlikely that the longer loops consist of 'pure' loops (i.e., all random coil or turn). In crystal structures of antibodies with medium to long CDR-H3 loops (McPC603 (Rudikoff et al., 1981): 11 amino acids (aa); KOL (Marquart et al., 1980): 17 aa; F19.9 (Lascombe et al., 1989): 15 aa) the loops consist of a disordered  $\beta$ -sheet extension from the  $\beta$ -barrel core and a 5-8 residue random coil/turn connecting these two strands.

To determine the nature of medium to long loops (>8 residues) which satisfy the CDR-H3 constraints, a complete search of the Protein Databank for loops of length 8-20 residues, was performed using the inter-C $\alpha$  distance constraints determined from known antibody crystal structures for CDR-H3. The resulting loops were then analyzed using the DSSP (Kabsch and Sander, 1983) program, which is able to assign secondary structure to polypeptide structures. The amount of secondary structure for each length of loop was calculated, and it was observed that for loops longer than 12 residues the amount of secondary structure within each of the classes described in DSSP was constant. The number of loops selected is also constant (approximately 150 loops) for loops longer than 12 residues. A closer inspection of each of the length ensembles shows indeed that the loops are the same between the groups.

This analysis shows that, like the long CDR-H3 crystal structures, the selected fragments consist of  $\beta$ -strands connected by 5-8 residue loops. For loops above 12-13 residues in length, the same loops are selected, but with extensions to the  $\beta$ -strands. This is called the "sliding-ladder" effect. In addition, the maximum size of a random coil or turn fragment in any of the structures contained in the Protein Databank tends not to exceed 8 residues, as determined by DSSP. This implies that the conformational space of longer loops is not saturated by the database and, although it is unlikely that long loops in antibodies will differ significantly from long loops in other structures, confidence in the prediction must be correspondingly reduced.

By how much is the usefulness of the CAMAL algorithm reduced by this observation?

The frequency of occurrence of different CDR-H3 lengths in antibody sequences described by Kabat et al. (Kabat et al., 1987) was analyzed. Figure 10 shows that more than 85% of H3 loops have lengths between 4 and 14 residues which can be modelled accurately by the CAMAL algorithm.

CDR-H3 of D1.3 is of average length (8 residues), though no loops of this length are seen in the available antibody structures. The crystal structure coordinate set showed an RMS of 1.9 Å compared with the model. The 36-71 loop is 12 residues long. The conformation is correctly predicted as a short loop connecting an

extension of the B-sheet.

The 3D6 H3 loop is 17 residues long. While KOL (Marquart et al., 1980) has the same length it has only one residue in common with 3D6 and only one conservative mutation. There is thus no reason to believe that the conformations would be similar. The final predicted conformation of 3D6 is an extended  $\beta$ -sheet, as in the crystal structure. The difference between the predicted and the crystal structure of 3D6-H3 is due to a twist of 5-7° in the extended  $\beta$ -sheet loop (see Figures 9A-9D). Such a twist has also been observed for complexed and uncomplexed antibodies by Wilson et al. (Wilson and others). This suggests that long CDR-H3 loops may be flexible and actively involved in antigen binding.

# 10 The Complete Variable Region

Prediction of the strand positions and  $V_L$ - $V_H$  orientation in the framework  $\beta$ -barrel was exact for all of the four antibodies. The backbone (N,C $\alpha$ ,C) RMS deviations from the crystal structures were between 0.56 and 0.86 Å, despite the fact that, in all cases the  $V_L$  and  $V_H$  regions of a particular model were derived from different antibody structures. This suggests that this method will do well in procedures such as humanization (Gorman et al., 1991), where correct framework positioning is important. The backbones of all six CDRs in all four antibodies are essentially correctly predicted, as shown in Figure 8. There are two important points to make about these predictions. First, the position of each CDR on its framework barrel is correct. Thus, CDR-framework interactions can be confidently monitored. The only deviation from the x-ray structure is CDR-H3 of antibody 3D6 which has been previously discussed. Second, the all atom RMS deviation between models and x-ray structures is dominated by sidechain positions. In most instances this deviation is due to a small number of incorrectly positioned, exposed sidechains (for example, in D1.3 the only sidechains which are incorrectly predicted are Tyr 9 of L1, Trp 4 of L3, Tyr 2 of H1 and Tyr 4 of H3). Since each CDR is constructed in the absence of other CDRs, the force field may choose a rotamer which is 120° away from that found in the crystal structure. This effect has also been observed by Lee et al. (Lee and Levitt, 1991).

#### Conclusion

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For antibodies having CDR H3 regions of 14 residues or less the complete variable domain can be modelled to an accuracy approaching that of medium resolution x-ray structures. For antibodies with longer H3 loops the CAMAL algorithm is likely to need an additional procedure in which molecular dynamics simulations are also incorporated.

The canonical approach of Chothia et al. appears to work well (at least in modelling backbones) where it may be applied and may be used successfully in combination with the CAMAL procedure.

One important observation that has emerged from these studies is that a given loop can exist in several conformations. In particular, this seems likely for CDR-L1 and, to a lesser extent, CDR-L3 and longer CDR-H3's. A simple combinatorial calculation shows that, if each of these three loops can exist in three separate conformations, a given combining site can have 3<sup>3</sup> = 27 different topographies. Clearly, this would explain the origins of cross reactivity and would allow for induced fit of antigens.

5	Table 2: Alignment of antibody sequences used in the modelling. '*' indicates (indicates β-strand regions used in the fitting for modelling frameworks. Nomencle strands is (H or L - Chain) - FR(Framework region)-(Strand number), thus for one of the heavy chain becomes HFR1.	gloop-2 d13 3671 3D6	Antibody gloop-2 dl3 3071	gloop-2 d13 3671 3D6	Antibody gloop-2 d13 3671 3D6
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35	ng. '*' in eworks. N number),	CHCC - 2	1777 - 7004 224 2004. 1-4-4.	+> Q>+- R+ + H H W D+ + + < < < < + · C + + < T < < + · C + + C C C C + · C + + C C C C + · C + + C C C C + · C +	WZIM+-
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			Framew	ork Model
Antibody	Resolution	R-factor	Light	Heavy
Gloop-2	2.80	21.2	REI	HyHEL-5
D1.3	-	-	REI	NEW
36-71	1.90	20.9	Gloop2	NEW
3D6	2.70	17.7	REI	KOL

Table 3: Details of the antibody crystal structures against which the models were compared and the parent frameworks used to build the models. Resolution data for D1.3 has not yet been published.

						RMS	(X) (X)					
	Antibody	CDR	sequence	SEQ ID NO	Co	N,Co,C All C	All CO	All MC	Ce	N,Co,C	VII CO	All MC
		-	BASIOISISISIYIS		9 7 9	9 71	2.01	<b>2</b>	9	0.07	2	2.13
	0.00	,	BACCONTONION		2 2 2		4.24	2	2.73	2.6	4.5	4.32
	36-71		RASIOCULINIFUN	98	2.71	2.43	•	4.84	3.61	9.31	5.5	8.67
	3D6		RAS Q (SIG) NINLH	497	0.81	0.84	2.48	1.02	0.81	0.78	2.00	1.2
	Glass-3	. 1	AASTI DS		0 38		5	8 .	•		1.10	1.10
	D1.3	į	ソープ・コープ・コープ		0.67	0.78		1.60	3	1.03	2.02	1.3
	36-71		PIT(SRS)OIS	300	0.0	0.00	2.24	2.23	e.7	0.73	3.6	2.40
	3D6		KASSLES	801	0.41	0.42	1.37	2	0.83	0.00	13	E
						}	}	<u>.</u>	:	:	}	3
	Gloop-3	L3	LQ[Y(LSY)P]LT	507	0.5	0.82	1.73					2
	34-71			6	2	100	2 2	2 10			2.37	2.28
	3D6		QIQ(YNS)YIS	808	<del>.</del>		3.04	3.86	2.31	1.97	3.00	9.96
												}
	Gloop-3	Ξ	[T(FGI)T]	306	0.8	0.70	2.8	1.60	1.04	1.01	2.04	2.00
	D1.3		[G(YGV)N]	107	0.4	0.83	2.31	2.08	3.0	0.30	3.26	2.90
	36-71		[S(NGI)N]	508	0.5	0.83	2.22	-	-	0.07	2.51	2.23
	3D6		DYAMH	509	0.67	0.77	1.63	E	0.81	0.73		1.70
	Gloop-2	3	EIFCPONSIKTY	810	2	•	=	1.70	<u>.</u>	2	2.28	2.10
	01.3		MIW(GDG)NITD	511	3	0.43	1.56		0.07	0.	-	1.00
	36-71		YNN[P(QNG)Y IA	813	0.0	0.78	2.01	2.20	1.47	1.11	1.73	1.90
	3D6		ISWDSSSIG	513	0.8	0.52	2.38	2.08	9	0.00	2.88	2.10
	Gleep-2	H <sub>2</sub>	IR(EIR)YI	\$ <b>1</b>	<u>.</u>	•	ب <u>د</u>	<b>3</b>	0.9	1.07	:	1.1
	D1.3		ERID(YRL)DIY	S1.5	<u>.</u>	0.53	-	- 2	1.2	9.2	ī	1.33
_	36-71		SEYY Q(QSY)K FDY	516	1.95	1.75	:	.08	2.65	2.83	8	.0
	3D6		GRDYY[D(SGG)YF]TVAPDI	517	3.66	3.42	5.93	4.01	4.30	3.96	9.30	6.30
_												

0.86 and 0.56 respectivly calculating the RMS over the loops. The total RMS of the frameworks (N,C $\alpha$ ,C) is 0.81, 0.60, calculated by least-squares fitting the conserved core of the two structures upon each other and difference between model and crystal structure loop coordinates. The RMS values are a global fit Table 4: Sequence and conformational search construction scheme for each of the 24 CDRs, =construction area, ( )= Chain closure, all sidechains are constructed. RMS(Root Mean Square) 5 15 20 25 30 35 45 50

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Loop	Canonical	Sequence	SEQ ID NO
Lļ	HyHEL-10	RASQSISRWLA	518
	(3D6)	RASQSIGNNLH	497
L2	REI	EASNDLA	519
	(3D6)	KASSLES	501
Hl	McPC603	DFYME	<b>520</b>
	(3D6)	DYAMH	50 <del>9</del>
H2	KOL	IIWDDGSDQ	521
	(3D6)	ISWDSSSIG	513

Table 5: Canonical loops selected for the model of 3D6(taken from Chothia et al (1989)).

an year	•		*				
Resi	due Number	24	25	26	27	28*	29*
ş=⊨ REI	Sequence	Q	Α	s	Q	S	ľ
	$\phi/\psi$	-/138	-103/157	-96/7	-158/142	-40/108	-112/9
HyHEL-10	Sequence	R	A	5	Q	S	I
	$\phi/\psi$	-/108	-85/135	-88/64	172/160	-64/-38	9/63
Resi	due Number	30*	31*	32	33	32	
REI	Sequence	I	К	Y	L	N	SEQ ID NO: 522
	$\phi/\psi$ .	79/-77	-146/21	-104/89	-143/133	-144/-	
HyHEL-10	Sequence	G	N	N	Ĺ	Н	SEQ ID NO: 518
-	$\phi/\psi$	-63/107	85/-15	-105/72	-129/118	-126/-	

Table 6: Backbone  $\phi$  and  $\psi$  angles of residues in CDR-L1 from HyHEL-10 and REI classified in the same canonical group by Chothia *et al* (1989). The residues exhibiting a peptide flip are indicated by a \*.

M.J. Darsley, P de al Paz, D.C. Phillips and A.R. Rees in Methodological Surveys in Biochemistry and Analysis, pages 63-68, Volume 15, 1985, Plenum Press (Eds. E. Reid, G.M.W. Cook and D.J. Morre), Presented at the Ninth International Subcellular Methodology Forum, September 3-6, 1984, Guildford, UK.

Amit, A.G., Mariuzza, R.A., Phillips, S.E.V. and Poljak, R.J. (1986). The Three-dimensional Structures of an Antigen-antibody Complex at 2.8 A Resolution. Science 233, pp. 747-753.

ాడీqvist, J., van Gunsieren, W., Leifonmark, M. and Tapia, O. (1985), J. Mol. Biol. 183, pp. 461-477.

Bernstein, F., Koetzle, T., Williams, G., Meyer, E., Brice, M., Rodgers, J., Kennard, O., Shimanouchi, T. and Tasumi, M. (1977), J. Mol. Biol. 112, pp. 535-542.

Brooks, B., Bruccoleri, R., Olaison, B., Statcs, D., Swaminathan, S. and Karplus, M. (1983), J. Comp. Chem. 4, pp. 187-217.

Bruccoleri, R.E. and Karplus, M. (1987), Prediction of the Folding of Short Polypeptide Segments by Uniform Conformational Sampling. Biopolymers 26, pp. 137-168.

Carter, D. et al. (1991). Protein Engineering, p. 9999.

Chau, P. and Dean P. (1987). Molecular Recognition: 3d Surface Structure Comparison by Gnomonic Projection. J. Mol. Graph. 5, pp. 97-100.

Chothia, C., Lesk, A., Levitt, M., Amit, A., Mariuzza, R., Phillips, S. and Poljak, R. (1986). The Predicted Structure of Immunoglobulin D1.3 and its Comparison with the Crystal Structure. Science 233, pp. 755-758.

Chothia, C., Lesk, A.M., Tramontano, A., Levitt, M., Smith-Gill, S.J., Air, G., Sheriff, S., Padlan, E.A., Davies, D.R., Tulip, W.R., Colman, P.M., Alzri, P.M. and Poljak, R.J. (1989). Conformations of Immunoglobulin Hypervariable Regions. Nature (London) 342, pp. 877-883.

Cornette, J.L., Cease, K.B., Margalit, H., Spouge, J.L., Berzofsky, J.A. and Delisi, C. (1987). Hydrophobicity Scales and Computational Techniques for Detecting Amphipatic Structures in Proteins. Journal of Molecular Biology 195.3, pp. 659-685.

Darsley, M. and Rees, A. (1985), EMBO J. 4, pp. 383-392.

Davies, D., Sheriff, S. and Padlan, E. (1988). Antibody Antigen Complexes. J. Biol. Chem. 263, pp. 10541-10544.

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de la Paz, P., Sutton, B., Darsly, M. and Rees, A. (1986). Modelling of the Combining Sites of Three Antilysozyme Monoclonal Antibodies and of the Complex Between One of the Antibodies and Its Epitope. EMBO J. 5, pp. 415-425.

Fine, R., Wang, H., Shenkin, P., Yarmush, D. and Levinthal, C. (1986). Predicting Antibody Hypervariable Loop Conformations ii: Minimization and Molecular Dynamics Studies of McPC603 from Many Randomly Generated Loop Conformations. Proteins: Struct., Funct., Genet. 1, pp. 342-362.

Go, N. and Sheraga, H. (1970). Ring Closure and Local Conformational Deformations of Chain Molecules. Macromolecules 3, pp. 178-187.

Gorman, S., Clark, M., Rutledge, E., Cobbold, S. and Waldman, H. (1991). Reshaping a Therapeutic CD4 Antibody. Proc. Natl. Acad. Sci. U.S.A. 88, pp. 4181-4185.

Grunow, R., Jahn, S., Porstman, T., Kiessig, T., Steinkeller, H., Steindl, F., Mattanovich, D., Gurtler, L., Deinhardt, F., Katinger, H. and von R., B. (1988). The High Efficiency, Human B Cell Immortalizing Heteromyeloma cb-f7. J. Immunol. Meth. 106, pp. 257-265.

Jeffrey, P. (1989). The Structure and Specificity of Immunoglobulins. D. Phil. Thesis, University of Oxford. Jeffrey, P.D., Gricst, R.E., Taylor, G.L. and Rees, A.R. (1991). Crystal Structure of the Fab Fragment of the Anti-peptide Antibody Gloop-2 and 2.8 Å. Manuscript in Preparation.

Kabat, E.A., Wu, T.T., Reid-Miller, M., Perry, B.M. and Gottesman, K.S. (1987). Sequences of Proteins of Immunological Interest. U.S. Department of Health and Human Services, Fourth Edition.

Kabsch, W. and Sander, C. (1983). Dictionary of Protein Secondary Structure. Biopolymers 22, pp. 2577-2637.

Lascombe, M., Alzari, P., Boulot, G., Salujian, P., Tougard, P., Berek, C., Haba, S., Rosen, E., Nisonof, A. and Poljak, R. (1989). Three-dimensional Structure of Fab r19.9, A Monoclonal Murine Antibody Specific for the p-azobenzenearsonate Group. Proc. Natl. Acad. Sci. U.S.A. **86**, p. 607.

Lee, C. and Levitt, M. (1991). Accurate Prediction of the Stability and Activity Effects of Site-directed Mutagenesis on a Protein Core. Nature **352.6334**, pp. 448-451.

Lee, C. and Subbiah, S. (1991). Prediction of Protein Side-chain Conformation by Packing Optimization. Journal of Molecular Biology **217.2**, pp. 373-388.

Marquart, M., Deisenhofer, J. and Huber, R. (1980), Crystallographic Refinement and Atomic Models of the Intact Immunoglobulin Molecule KOL and Its Antigen-binding Fragment at 3.0 Å and and Resolution. J. Mol. Biol. 141, pp. 369-391.

Martin, A.C.R. (1990). Molecular Modelling of Antibody Combining Sites. D. Phil. Thesis, University of Oxford.

Martin, A.C.R., Cheetham, J.C. and Rees, A.R. (1989). Modelling Antibody Hypervariable Loops: A Combined Algorithm. Proc. Natl. Acad. Sci. U.S.A. 86, pp. 9268-9272.

Martin, A.C.R., Cheetham, J.C. and Rees, A.R. (1991). Modelling Antibody Hypervariable Loops using a 'Combined Algorithm'. Meth. Enz. In press.

Moult, J. and James, N. (1986). Proteins: Struct., Funct., Genet. 1, p. 146.

Padlan, E., Davies, D., Pecht, I., Givol, D. and Wright, C. (1976). Model Building Studies of Antigen-binding Sites: The Hapten-Binding Site of MOPC-315. Cold Spring Harbor Quant. Symp. Biochem. 41, pp. 627-637.

Palmer, K. and Sheraga, J. (1991). Standard-geometry Chains Fitted to X-ray Deviated Structures: Validation of the Rigid-geometry Approximation. I. Chain Closure through a Limited Search of Loop Conformations. J. Comp. Chem. 12, pp. 505-526.

Paul, P., Burney, P., Campbell, M. and Odguthorpe, D. (1990). The Conformational Preferences of γ-lactam and Its Role in Constraining Peptide Structure. J. Comp.-aided. Mol. Des. 4, pp. 239-253.

Rose, D.R., Strong, R.K., Margolis, M.N., Gefter, M.L. and Petsko, G.A. (1990). Crystal Structure of th Antigen-binding Fragment of the Murine Anti-arsonate Monoclonal Antibody 36-71 at 2.9 Å Resolution. Proc. Natl. Acad. Sci. U.S.A. 87, pp. 338-342.

Rudikoff, S., Satow, Y., Padlan, E.A., Davies, D.R. and Potter, M. (1981). Kappa Chain Structure from a Crystallized Murine Fab': The Role of the Joining Segment in Hapten Binding. Mol. Immunol. 18, pp. 705-711.

Schilling, J., Clevinger, B., Davie, J.M. and Hood, L. (1980). Amino Acid Sequence of Homogeneous Antibodies to Dextran and DNA Rearrangements in Heavy Chain V-region Gene Segments. Nature (London) 283, pp. 35-40.

Strong, R., Campbell, R., Rose, D., Petsko, G., Sharon, J. and Margolies, M. (1991). Three-dimensional Structure of Murine Anti-p-azophenylarsonate Fab 36-71.1, X-ray Crystallography, Site-directed Mutagenesis, and Modeling of the Complex with Hapten. Biochemistry **30**, pp. 3739-3748.

Thornton, J., Sibanda, B., Edwards, M. and Barlow, D. (1988). Analysis, Design and Modification of Loop Regions in Proteins. BioEssays 8, pp. 63-69.

Tramontano, A. Chothia, C. and Lesk, A. (1989). Structural Determinants of the Conformations of Medium-

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sized Loops in Proteins. Proteins: Struct., Funct., Genet. 6, pp. 382-394. Wilson, I. et al., Presented at Structure and Function Meeting in Honour of Sir David Phillips, 1-3 July, 1991, Oxford, UK. SEQUENCE LISTING GENERAL INFORMATION (i) APPLICANT: PEDERSEN, Jan T. SEARLE, Stephen M.J. Anthony R. REES. ROGUSKA, Michael A. GUILD, Braydon C. (ii) TITLE OF INVENTION: SURFACE RESIDUE VENEERING OF RODENT ANTIBODIES (iii) NUMBER OF SEQUENCES: 522 (iv) CORRESPONDENCE ADDRESS: (A) ADDRESSEE: Sughrue, Mion, Zinn, Macpeak & Seas (B) STREET: 2100 Pensylvania Avenue, N.W. (C) CITY: Washington (D) STATE: D.C. (E) COUNTRY: United States (F) ZIP: 20037-3202 (v) COMPUTER READABLE FORM: (A) MEDIUM TYPE: Floppy disk (B) COMPUTER: HP 9000/700 Workstation (C) OPERATING SYSTEM: UNIX (D) SOFTWARE: In house (vi) CURRENT APPLICATION DATA: (A) APPLICATION NUMBER: 07/942,245 (B) FILING DATE: 09-SEP-1992 (C) CLASSIFICATION: (ix) TELECOMMUNICATION INFORMATION: (A) TELEPHONE: (202) 293-7060 (B) TELEFAX: (202) 293-7860 (C) TELEX: 6491103 (1) INFORMATION FOR SEQ ID NO:1 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 109 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:1:

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Asp Ile Gln Met Thr Gln Ser Pro Ser Ser Leu Ser Ala Ser L u Gly

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5	Glu	Arg	Val	Ser 20	Leu	Thr	Cys	Arg	Ala 2	Ser 5	Gln	Glu	Ile		Gly 30	Tyr
10	Leu	Ser	Trp 35	Leu	Gln	Gln	Lys	Pro 40	Asp O	Gly	Thr	Ile		Arg 15	Leu	Ile
	Tyr	Ala 50	Ala	Ser	Thr	Leu	Asp 55	Ser	Gly	Val	Pro		Arg 0	Phe	Ser	Gly
15	Arg 65	Arg	Ser	Gly	Ser	Asp 70	Tyr	Ser	Leu	Thr	11e 75	Ser	Ser	Leu	Glu	Ser 80
	Glu	Asp	Phe	Ala	Asp 85	туг	Tyr	Cys	Leu	Gln 90	Tyr O	Leu	Ser	Tyr	_	Leu 5
20	Thr	Phe	Gly	Ala 100		Thr	Lys	Leu	Glu 105		Lys	s Arq	, Ala	a		
25	(2)			ION SEQU					TICS	3 <b>:</b>						
				(1	A) L B) T	ENGT YPE:	H: 1 ami OGY:	09 a no a	mino cid		.ds					
30				MOLE SEQU				_		, 'O TD	NO.					
35	Asp 1	Ile				Gln					Leu		Ala	Ser		Gly L5
	Glu	Thr	Val	Thr 20	Ile	Thr	Cys	Arg	Ala 25		Gly	Asn	Ile		Asn 0	Tyr
40	Leu	Ala_	Trp 35	Tyr	Gln	Gln	Lys	Gln 40		Lys	Ser	Pro	_	Leu 5	Leu	Val
<b>4</b> 5	Tyr	Tyr 50	Thr	Thr	Thr	Leu	Ala 55		Gly	Val	Pro	Ser 6	_	Phe	Ser	Gly
50	Ser 65	Gly	Ser	Gly	Thr	Gln 70	Tyr	Ser	Leu	Lys	Ile 75	Asn	Ser	Leu	Gln	Pro 80
	Glu	Asp	Phe	Gly	Ser 85	Tyr	Tyr	Cys	Gln	His 90		Trp	Ser	Thr	_	Arg 5

Thr Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys Arg Arg 100 105

(3) INFORMATION FOR SEQ ID NO:3

10			(1)	(1	A) L	ENGT YPE:	H: 1 ami	.07 a	mino cid		.ds					
		(	ii)	MOLE	CULE	TYP	E: p	epti	de							
15		(	xi)	SEQU	ENCE	DES	CRIP	TION	: SE	Q II	) NO:	3:				
	Asp 1	Ile	Val	Leu	Thr 5	Gln	Ser	Pro	Ala	Ile 1		Ser	Ala	Ser		Gly 15
20	Glu	Lys	Val	Thr 20	Met	Thr	Cys	Ser	Ala 25		Ser	Ser	Val		Tyr 0	Met
25	Tyr	Trp	Tyr 35	Gln	Gln	Lys	Ser	Gly 40		Ser	Pro	Lys	_	Trp 5	Ile	туг
	Asp	Thr 50	Ser	Lys	Leu	Ala	Ser 55		Val	Pro	Val		Phe 0	Ser	Gly	Ser
30	Gly 65		Gly	Thr	Ser	Tyr 70	Ser	Leu	Thr	Ile	Ser 75	Ser	Met	Glu	Thr	Glu 80
35	Asp	Ala	Ala	Glu	Tyr 85	Tyr	Cys	Gln	Gln	Trp 9		Arg	Asn	Pro		Phe 5
	Gly	Gly	Gly	Thr 100	Lys	Leu	Glu	Ile	Lys 105		j Ala	<b>a</b> .				
40	<b>(-4</b> )	INFO	RMAT	ION :	FOR	SEQ	ID N	io: 4								
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50	Asp 1	Ile	Val	Leu	Thr 5		Ser	Pro	Ala	Thr 1		Ser	Val	Thr		Gly 15

5	Asn	Ser	Val	Ser 20	Leu	Ser	Cys	Arg	Ala 25		Gln	Ser	Ile		Asn 0	Asn
	Leu	His	Trp 35	Tyr	Gln	Gln	Lys	Ser 40		Glu	Ser	Pro	_	Leu 5	Leu	Ile
10	Lys	Tyr 50	Ala	Ser	Gln	Ser	Ile 55		Gly	Ile	Pro		Arg O	Phe	Ser	Gly
15	Ser 65	Gly	Ser	Gly	Thr	Asp 70	Phe	Thr	Leu	Ser	Ile 75	Asn	Ser	Val	Glu	Thr 80
	Glu	Asp	Phe	Gly	Met 85	Tyr	Phe	Cys	Gln	Gln 9	_	Asn	Ser	Trp	Pro 9	Tyr 5
20	Thr	Phe	Gly	Gly 100	Gly	Thr	Lys	Leu	Glu 105		Lys	s Ar	g Ala	a ·		
	(5)	[NFO	RMAT	ION :	FOR	SEQ	ID N	0:5								
25			(i) :	(	A) L	ENGT	H: 1		TICS		ids					
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30		(:	ii) I	į,	C) T	OPOL	OGY:	lir	ear							
30		Ť		() MOLE	CULE	OPOL TYP	OGY: E: p	lir epti	ear	EQ II	о ио:	:5:				
30 35	Glu 1	(:	xi)	() MOLE SEQU	c) t cule ence	OPOL TYP DES Gln	OGY: E: p CRIP	lir epti TION	near .de J: SI	Ile			Ala	Ser	<b>Leu</b>	Gly L5
	1	(:	xi) : Val	() MOLE SEQU Leu	C) TO CULE ENCE Thr 5	TYP DES Gln	OGY: E: p CRIF Ser	lir epti TION Pro	near .de J: SE Ala	Ile 1	Thr O	Ala		Ser		L5
	1 Gln	(; Ile Lys	xi) (	MOLE SEQU Leu Thr 20	C) TO CULE ENCE Thr 5	TYP DES Gln	OGY: E: p CRIP Ser Cys	lir epti PTION Pro Ser	de J: SE Ala Ala 2	Ile 1 Ser 5	Thr 0 Ser	Ala	Val Pro	Ser	Ser	Leu
35	1 Gln His	Ile Lys	Val Val Tyr 35	MOLE SEQU Leu Thr 20	C) TO CULE ENCE Thr 5	OPOL TYP DES Gln Thr	OGY: E: p CRIF Ser Cys	lir epti PTION Pro Ser Gly 4	de U: SE Ala Ala 2: Thr	Ile 1 Ser 5	Thr 0 Ser Pro	Ala Ser Lys	Val Pro	Ser Trp	Ser 30	Leu Tyr
35 40	Gln His Glu	Lys Trp	Val Val Tyr 35	MOLE SEQU Leu Thr 20 Gln	C) TO CULE ENCE Thr 5 Ile Gln Leu	OPOL TYP DES Gln Thr Lys	OGY: E: p CRIF Ser Cys Ser	lir epti PTION Pro Ser Gly Gly	de U: SE Ala Ala 2: Thr O	Ile 1 Ser 5	Thr 0 Ser Pro	Ala Ser Lys Arg	Val Pro Phe	Ser Trp 15	Ser 30	Leu Tyr Ser

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10	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 112 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:
	Glu Ser Val Leu Thr Gln Pro Pro Ser Ala Ser Gly Thr Pro Gly Gln 1 5 10 15
20	Arg Val Thr Ile Ser Cys Thr Gly Thr Ser Ser Asn Ile Gly Ser Ile 20 25 30
25	Thr Val Asn Trp Tyr Gln Gln Leu Pro Gly Met Ala Pro Lys Leu Leu 35 40 45
30	Ile Tyr Arg Asp Ala Met Arg Pro Ser Gly Val Pro Thr Arg Phe Ser 50 60
30	Gly Ser Lys Ser Gly Thr Ser Ala Ser Leu Ala Ile Ser Gly Leu Glu 65 70 75 80
35	Ala Glu Asp Glu Ser Asp Tyr Tyr Cys Ala Ser Trp Asn Ser Ser Asp 85 90 95
	Asn Ser Tyr Val Phe Gly Thr Gly Thr Lys Val Thr Val Leu Gly Gln 100 105 110
40	(7) INFORMATION FOR SEQ ID NO:7  (i) SEQUENCE CHARACTERISTICS:
45	(A) LENGTH: 115 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:7:
50	Asp Ile Val Met Thr Gln Ser Pro Ser Ser Leu Ser Val Ser Ala Gly 1 5 10 15
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5	Glu	Arg	Val	Thr 20	Met	Ser	Cys	Lys	Ser 25	Ser 5	Gln-	Ser	Leu		Asn 0	Ser
	Gly	Asn	Gln 35	Lys	Asn	Phe	Leu	Ala 40		Tyr	Gln	Gln		Pro 5	Gly	Gln
10	Pro	Pro 50	Lys	Leu	Leu	Ile	<b>Tyr</b> 55	Gly	Ala	Ser	Thr		Glu 0	Ser	Gly	Val
15	Pro 65	Asp	Arg	Phe	Thr	Gly 70	Ser	Gly	Ser	Gly	Thr 75	Asp	Phe	Thr	Leu	Thr 80
	Ile	Ser	Ser	Val	Gln 85	Ala	Glu	Asp	Leu	Ala 90		Tyr	Tyr	Сув		Asn 5
20	Asp	His	Ser	Tyr 100	Pro	Leu	Thr	Phe	Gly 10		Gly	Thr	Lys	Leu 11		Ile
25	Lys	Arg	Ala 115													
	(8)	INFO	RMAT:	ION :	FOR	SEQ	ID N	0:8								
<b>30</b>			(i) :	(,	A) L B) T	ENGT YPE:		03 a	mino cid		ds					
		(:	ii) i	MOLE	CULE	TYP	E: p	epti	de							
35		(:	xi)	SEQU	ENCE	DES	CRIP	TION	: SE	Q II	) NO:	8:				
	Ser 1	Val	Leu	Thr	Gln 5		Pro	Ser	Val	Ser 1	_	Ala	Pro	Gly		Arg 15
40	Val	Thr_	Ilē	Ser 20	Cys	Thr	Gly	Ser	Ser 25		Asn	Ile	Gly		Gly 0	Asn
<b>4</b> 5	His	Val	Lys 35	Trp	Tyr	Gln	Gln	Leu 40	_	Gly	Thr	Ala		Lys 5	Leu	Leu
50	İle	Phe 50	His	Asn	Asn	Ala	Arg 55		Ser	Val	Ser		Ser 0	Gly	Ser	Ser
	Ala 65	Thr	Leu	Ala	Ile	Thr 70	Gly	Leu	Gln	Ala	<b>Glu</b> 75	Asp	Glu	Ala	Asp	Tyr 80

5	Tyr	Cys	Gln	Ser	Tyr 85	Asp	Arg	Ser	Leu	Arg 90		Phe	Gly	Gly	_	Thr 5
	Lys	Leu	Thr	Val 100		Arg	Gln	1	-							
10	(9)	INFO	RMAT	ION	FOR	SEQ	ID N	0:9								
15			(i)	(	A) L B) T	ENGT YPE:	H: 1	.14 a	minc cid	3: o aci	ds					
		(	ii)	MOLE	CULE	TYP	E: p	epti	.de							
		. (:	xi)	SEQU	ENCE	DES	CRIP	TION	l: SE	Q ID	NO:	9:				
20	Asp 1	Val	Val	Met	Thr 5		Thr	Pro	Leu	Ser 10		Pro	Val	Ser		Gly .5
25	Asp	Gln	Ala	Ser 20	Ile	Ser	Cys	Arg	Ser 25		Gln	Ser	Leu		His O	Ser
	Gln	Gly	Asn 35	Thr	Tyr	Leu	Arg	Trp		Leu	Gln	Lys		Gly 5	Gln	Ser
30	Pro	Lys 50	Val	Leu	Ile	Tyr	Lys 55		Ser	Asn	Arg		Ser 0	Gly	Val	Pro
35	Asn 65	Arg	Phe	Ser	Gly	Ser 70	Gly	Ser	Gly	Thr	Asp 75	Phe	Thr	Leu	Lys	Ile 80
40	Ser	Arg	Val	Glu	Ala 85	Glu	Asp	Leu	Gly	Val 90		Phe	Cys	Ser	_	Ser 5
	Thr	His	_Val	Pro 100	Trp	Thr	Phe	Gly	Gly 10		Thr	Lys	Leu	Glu 11		Lys
<b>4</b> 5	Arg	Ala														
	(10)	INF	ORMA	TION	FOR	SEQ	ID	NO: 1	LO		٠					
50			(i)	(		ENGT YPE:	H: 1 ami	.09 a	mino acid	s: aci	ds		,			
		(	ii)	MOLE	CULE	TYP	E: p	epti	ide							
55																

		(	xi)	SEQU	ENCE	DES	CRIP	TION	: SE	EQ II	ON C	:10:				
5	Asp 1	Ile	Gln	Met	Thr 5	Gln	Thr	Thr	Ser	Ser 1	Leu 0	Ser	Ala	Ser		Gly 15
10	Asp	Arg	Val	Thr 20	Ile	Ser	Cys	Arg	Ala 25	Ser	Gln	Asp	Ile		Asn 0	Tyr
15	Leu	Asn	Trp 35	Tyr	Gln	Gln	Lys	Pro 40	Asp )	Gly	Thr	Val		Leu 5	Leu	Val
,,	Tyr	Tyr 50	Thr	Ser	Arg	Leu	His 55	Ser	Gly	Val	Pro		Arg 0	Phe	Ser	Gly
20	Ser 65	Gly	Ser	Gly	Thr	Asp 70	туг	Ser	Leu	Thr	Ile 75	Ser	Asn	Leu	Glu	His 80
	Glu	Asp	Ile	Ala	Thr 85	Tyr	Phe	Cys	Gln	Gln 90		Ser	Thr	Thr	_	Arg 5
?5	Thr	Phe	Gly	Gly 100	Gly	Thr	Lys	Leu	Glu 105		. Lys	s Ar	g Arq	3		
30	(11)			SEQUI () (1)	ENCE A) LI B) T	CHA ENGT: YPE:	RACT	ERIS 09 a no a	TICS mino cid		.ds					
35				MOLE				•								
10	Asp 1			SEQUI Met		Gln					Leu		Ala	Ser		Gly .5
	Asp	Arg	Val	Ser 20		Ser	Cys	Arg	Ala 25		Gln	Asp	Ile	Asn 3	Asn O	Phe
15	Leu	Asn	Trp 35	Tyr	Gln	Gln	Lys	Pro 40		Gly	Thr	Ile		Leu 5	Leu	Ile
ю	Tyr	Phe 50	Thr	Ser	Arg	Ser	Gln 55		Gly	Val	Pro		Arg 0	Phe	Ser	Glý

5	,	75
	Glu	Asp Ile Ala Thr Tyr Phe Cys Gln Gln Gly Asn Ala Leu Pro Arg 85 90 95
10	Thr	Phe Gly Gly Gly Thr Lys Leu Glu Ile Lys Arg Ala
	(12)	INFORMATION FOR SEQ ID NO:12
15		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 107 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
20		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:12:
25	Asp 1	Ile Gin Met Thr Gln Ser Pro Ser Thr Leu Ser Ala Ser Val Gly 5 10 15
	Asp	Arg Val Thr Ile Thr Cys Arg Ala Ser Gln Ser Ile Ser Arg Trp 20 25 30
30	Leu	Ala Trp Tyr Gln Gln Lys Pro Gly Lys Val Pro Lys Leu Leu Ile 35 40 45
35	Tyr	Lys Ala Ser Ser Leu Glu Ser Gly Val Pro Ser Arg Phe Ser Gly 50 55 60
	Ser 65	Gly Ser Gly Thr Glu Phe Thr Leu Thr Ile Ser Ser Leu Gln Pro
40	Asp	Asp Phe Ala Thr Tyr Tyr Cys Gln Gln Tyr Asn Ser Tyr Ser Phe 85 90 95
45	Gly	Pro Gly Thr Lys Val Asp Ile Lys Arg Thr 100 105
,,,	(13)	INFORMATION FOR SEQ ID NO:13
50		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 104 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide

42

		(	xi)	SEQU	ENCE	DES	CRIP	TION	ı: SE	Q II	NO:	:13:				
5	Gln 1	Val	Gln	Leu	Gln 5	Gln	Ser	Gly	Thr	Glu 1	Leu 0	Ala	Arg	Pro		Ala L5
10	Ser	Val	Arg	Leu 20	Ser	Cys	Lys	Ala	Ser 25		Tyr	Thr	Phe	_	Thr 0	Phe
15	Gly	Ile	Thr 35	Trp	Val	Lys	Gln	Arg 40		Gly	Gln	Gly	_	Glu 5	Trp	Ile
	Gly	Glu 50	Ile	Phe	Pro	Gly	Asn 55	Ser	Lys	Thr	Tyr		Ala O	Glu	Arg	Phe
20	Lys 65	Gly	Lys	Ala	Thr	Leu 70	Thr	Ala	Asp	Lys	Ser 75	Ser	Thr	Thr	Ala	Tyr 80
	Met	Gln	Leu	Ser	Ser 85	Leu	Thr	Ser	Glu	Asp 90		Ala	Val	Tyr	_	Cys 5
25	Ala	Arg	Glu	Ile 100	Arg	Tyr	Trp	Gly	•							
30	(14)			(1	ENCE A) L B) T	CHA ENGT YPE:	RACT	ERIS 07 a no a	TICS mino	: aci	.ds					
35				MOLE SEQU				_		o .TD	NO:	14:				
<b>4</b> 0	Gln 1	•		Leu		Glu			,		Leu		Ala	Pro		Gln IS
	Ser	Leu	Ser	Ile 20	Thr	Cys	Thr	Val	Ser 25	_	Phe	Ser	Leu	_	Gly O	Tyr
<b>4</b> 5	Gly	Val	Asn 35	Trp	Val	Arg	Gln	Pro 40		Gly	Lys	Gly		Glu 5	Trp	Leu
50	Gly	Met 50	Ile	Trp	Gly	Asp	<b>Gly</b> 55		Thr	Asp	Tyr		Ser 0	Ala	Leu	Lys

5	Ser 65	Arg	Leu	Ser	Ile	Ser 70	Lys	Asp	Asn	Ser	Lys 75	Ser	Gln	Val	Phe	Leu 80
	Lys	Met	Asn	Ser	Leu 85	His	Thr	Asp	qsA	Thr 90		Arg	Tyr	Tyr		Ala 5
10	Arg	Glu	Arg	Asp 100	Tyr	Arg	Leu	Asp	Tyr 105	_	Gly	7				
	(15)	INF	ORMA	TION	FOR	SEQ	ID	NO:1	5							
15	<sup></sup>		(i)	(1	ENCE A) L B) T C) T	ENGT YPE:	H: 1 ami	06 a no a	mino cid		.ds					
		(	ii)	MOLE	CULE	TYP	E: p	epti	de							
20		(:	xi)	S <b>EQ</b> U	ENCE	DES	CRIP	TION	: SE	Q II	NO:	15:				
	Val 1	Gln	Leu	Gln	Gln 5		Gly	Ala	Glu	Leu 1		Lys	Pro	Gly		Ser 15
25	Val	Lys	Ile	Ser 20	Cys	Lys	Ala	Ser	Gly 25	_	Thr	Phe	Ser	_	Tyr 0	Trp
30	Ile	Glu	Trp 35	Val	Lys	Gln	Arg	Pro 40		His	Gly	Leu		Trp 5	Ile	Gly
35	Glu	Ile 50		Pro	Gly	Ser	Gly 55		Thr	Asn	Tyr	_	Glu 0	Arg	Phe	Lys
	Gly 65	Lys	Ala	Thr	Phe	Thr 70	Ala	Asp	Thr	Ser	Ser 75	Ser	Thr	Ala	Tyr	Met 80
40	Gln	Leu	Asn 	Ser	Leu 85		Ser	Glu	Asp	Ser 9	_	Val	Tyr	Tyr		Leu 95
45	His	Gly	Asn	Tyr 100		Phe	Asp	Gly	Trp 105		7					
40	(16)	INF	ORMA	TION	FOR	SEQ	ID	NO: 1	.6							
50			(i)	į	ENCE A) L B) T C) T	ENGT	H: 1 ami	.04 a	mind		ids					
		(	ii)	MOLE	•											

		(:	xi)	SEQU	ENCE	DES	CRIP	TION	: SE	Q IE	NO:	:16:				
5	Asp 1	Val	Gln	Leu	Gln 5	Glu	Ser	Gly	Pro	Ser 1		Val	Lys	Pro		Gln L5
10	Thr	Leu	Ser	Leu 20	Thr	Cys	Ser	Val	Thr 25		Asp	Ser	Ile	_	Ser 0	Asp
15	Tyr	Trp	Ser 35	Trp	Ile	Arg	Lys	Phe 40		Gly	Asn	Arg		Glu 5	Tyr	Met
15	Gly	Tyr 50	Val	Ser	Tyr	Ser	Gly 55	Ser	Thr	Tyr	Tyr		Pro 0	Ser	Leu	Lys
20	Ser 65	Arg	Ile	Ser	Ile	Thr 70	Arg	Asp	Thr	Ser	Lys 75	Asn	Gln	Tyr	Tyr	Leu 80
25	Asp	Leu	Asn	Ser	Val 85	Thr	Thr	Glu	Asp	Thr 90		Thr	Tyr	Tyr		Ala 95
	Asn	Trp	Asp	Gly 100	Asp	Tyr	Trp	Gly	•							
30	(17)			(1	ENCE A) Li B) T	CHA ENGT YPE:	RACT H: 1 ami	ERIS	TICS mino cid		ds					
35				MOLE			_	_		10 TE						
40	Glu 1			SEQUI Leu		Glu					Leu		Gln	Pro		Gly 15
	Ser	Leu	Lys	Leu 20	Ser	Cys	Ala	Ala	Ser 25	_	Phe	Asp	Phe	_	Lys 0	Tyr
<b>4</b> 5	Trp	Met	Ser 35	Trp	Val	Arg	Gln	Ala 40		Gly	Lys	Gly		Glu 5	Trp	Ile
50	Gly	Glu 50	Ile	His	Pro	Àsр	Ser 55	_	Thr	Ile	Asn		Thr 0	Pro	Ser	Leu

45

5	•			Phe		70					75					80
	Leu	Gln	Met	Ser	Lys 85	Val	Arg	Ser	Glu	Asp 9	Thr O	Ala	Leu	Tyr		Cys 95
10	Ala	Arg	Leu	His 100	Tyr	Tyr	Gly	Туг	Asn 105	Ala	а Туз	Tr	o Gl	Y		
	(18)	INF	ORMA	TION	FOR	SEQ	ID	NO: 1	.8							
15	.X.		(i)	(1	A) L B) T	ENGT YPE:	RACT H: 1 ami OGY:	.17 a	mino cid		ds					
20		(	ii)	MOLE	CULE	TYP	E: p	epti	de							
		(	xi)	SEQU	ENCE	DES	CRIP	TION	: SE	Q II	NO:	18:				
25	Glu 1	Val	Gln	Leu	<b>Val</b> 5	Gln	ser	Gly	Gly	Gly 10		Val	Gln	Pro	_	Arg L5
	Ser	Leu	Arg	Leu 20	Ser	Cys	Ser	Ser	Ser 25		Phe	Ile	Phe		Ser 0	Tyr
30	Ala	Met	Tyr 35	Trp	Val	Arg	Gln	Ala 40		Gly	Lys	Gly	_	Glu 5	Trp	Val
35	Ala	Il <b>e</b> 50	Ile	Trp	Asp	Asp	Gly 55	Ser	Asp	Gln	His	Tyr 6		Asp	Ser	Val
	Lys 65	Gly	Arg	Phe	Thr	Ile 70	Ser	Arg	Asn	Asp	Ser 75	Lys	Asn	Thr	Leu	Phe 80
40	Leu	Gln	Met -	Asp	Ser 85	Leu	Arg	Pro	Glu	Asp 90		Gly	Val	Tyr		Cys 5
<b>4</b> 5	Ala	Arg	Asp	Gly 100	Gly	His	Gly	Phe	Cys 105		Ser	Ala	Ser	Cys 11		Gly
	Pro	Asp	Tyr 115	Trp	Gly											
50	(19)	INF	ORMA!	PION	FOR	SEQ	ID 1	NO:1	9					•		
			(i) :		A) L	engt:	H: 1	13 a	mino		ds					
55				1)	o) T	ir£;	ami	no a	CIQ							

				,	<b>C)</b> 1	OPUL	JUGY :	11r	near							
5		(	ii)	MOLE	CULE	TYF	E: p	epti	de							
		(	xi)	SEQU	ENCE	DES	CRIE	MOIT	ı: sı	EQ II	ON C	:19:				
10	Glu 1	Val	Lys	Leu	Val 5	Glu	Ser	Gly	Gly	Gly 1		Val	Gln	Pro	_	Gl <sub>3</sub> 15
	Ser	Leu	Arg	Leu 20	Ser	Сув	Ala	Thr	Ser 29	Gly 5	Phe	Thr	Phe	_	Asp 80	Phe
15	Tyr	Met	Glu 35	Trp	Val	Arg	Gln	Pro	Pro	Gly	Lys	Arg		Glu 5	Trp	Ile
20	Ala	Ala 50	Ser	Arg	Asn	Lys	Gly 55	Asn	Lys	Tyr	Thr		Glu 0	туг	Ser	Ala
25	Ser 65	Val	Lys	Gly	Arg	Phe 70	Ile	Val	Ser	Arg	Asp 75	Thr	Ser	Gln	Ser	Ile 80
	Leu	Tyr	Leu	Gln	Met 85	Asn	Ala	Leu	Arg	Ala 90		Asp	Thr	Ala		Tyr 95
30	Туг	Cys	Ala	Arg 100	Asn	Tyr	Tyr	Gly	Ser 10		Trp	Tyr	Phe	Asp 11	_	Trp
35	Gly															
	(20)	INF	ORMA'	TION	FOR	SEQ	ID	NO: 2	0							
40		-	(i) :	(1	A) L: B) T	engt Ype:		07 a no a	mino cid	: aci	.ds					
		(:	ii) 1	MOLE	CULE	TYP	E: p	epti	de							
45		(2	Ki)	SEQU	ENCE	DES	CRIP	TION	: SE	Q ID	NO:	20:				
	Val 1	Gln	Leu	Glu	Gln 5		Gly	Pro	Gly	Leu 1		Arg	Pro	Ser		Thr 15
50	Leu	Ser	Leu	Thr 20	Cys	Thr	Val	Ser	Gly 25		Ser	Phe	Asp	_	Tyr 0	Tyr
											-				•	

5	Ser	Thr	Trp 35	Val	Arg	Gln	Pro	Pro 40	Gly <sup>.</sup>	Arg	Gly	Leu		Trp 5	Ile	Gly
	Tyr	<b>Val</b> 50	Phe	Tyr	His	Gly	Thr 55	Ser	Asp	Thr	Asp		Pro 0	Leu	Arg	Ser
10	Arg 65	Val	Thr	Met	Leu	Val 70	Asn	Thr	Ser	Lys	Asn 75	Gln	Phe	Ser	Leu	Arg 80
15		Ser	Ser	Val	Thr 85	Ala	Ala	Asp	Thr	Ala 9		Tyr	Tyr	Cys	_	Arg 5
	Asn	Leu	Ile	Ala 100	Gly	Cys	Ile	asp	Val 105	_	Gly	?				
20	(21)	INF	ORMA'	rion	FOR	SEQ	ID	NO:2	1							
25			(i) :	(1	A) L B) T	ENGT YPE:	H: 1 ami	ERIS 09 a no a lin	mino cid		ids					
		(:	ii) 1	MOLE	CULE	TYP	E: p	epti	de			·				
-		(:	xi) :	SEQUI	ENCE	DES	CRIP	TION	: SE	Q II	NO:	21:				
30	Glu 1	Val	Lys	Leu	<b>Asp</b> 5		Thr	Gly	Gly	Gly 1		Val	Gln	Pro		Arg .5
35	Pro	Met	Lys	Leu 20	Ser	Cys	Val	Ala	Ser 25		Phe	Thr	Phe	_	Asp 0	Tyr
	Trp	Met	Asn 35	Trp	Val	Arg	Gln	Ser 40		Glu	Lys	Gly		Glu 5	Trp	Val
40	Ala	Gln 50		Arg	Asn	Lys	Pro 55	_	Asn	Tyr	Glu	_	Tyr 0	Tyr	Ser	Asp
<b>45</b>	Ser 65	Val	Lys	Gly	Arg	Phe 70	Thr	Ile	Ser	Arg	<b>Asp</b> 75	Asp	Ser	Lys	Ser	Ser 80
50 -	Val	Tyr	Leu	Gln	Met 85		Asn	Leu	Arg	Val 9		Asp	Met	Gly	_	Tyr 95
<i></i>	Tyr	Cys	Thr	Gly 100	Ser	Tyr	Tyr	Gly	Met 105		э Ту	r Tr	p Gl	Y		
55	(22)	INF	ORMA	TION	FOR	SEQ	ID	NO:2	22							_

5			(1)	(	A) L B) T		H: 1 ami	.15 a	mino		ds					
10				MOLE SEQU						Q IC	NO:	:22:				
	Gln 1	Val	Gln	Leu	Lys 5	Glu	Ser	Gly	Ala	Glu 1		Val	Ala	Ala		Ser 15
15	Ser	Val	Lys	Met 20	Ser	Cys	Lys	Ala	Ser 25		Tyr	Thr	Phe		Ser 0	Tyr
20	Gly	Val	Asn 35	Trp	Val	Lys	Gln	Arg 40	Pro	Gly	Gln	Gly		Glu 5	Trp	Ile
25	Gly	Tyr 50	Ile	Asn	Pro	Gly	Lys 55		Tyr	Leu	ser		Asn O	Glu	Lys	Phe
	Lys 65	Gly	Lys	Thr	Thr	Leu 70	Thr	Val	Asp	Arg	Ser 75	Ser	Ser	Thr	Ala	Tyr 80
30	Met	Gln	Leu	Arg	Ser 85		Thr	Ser	Glu	Asp 90		Ala	Val	Tyr		Cys 5
35	Ala	Arg	Ser	Phe 100	Tyr	Gly	Gly	Ser	Asp 105		Ala	Val	Tyr	Tyr 11		Asp
	Ser	Trp	Gly 115													
40	(23)	inf	orha	TION	FOR	SEQ	ID	ио: 3	3							
			<u>(</u> i) <u> </u>	į.	A) L B) T		H: 1 ami	12 a	minc cid		ds					
45		(	ii)	MOLE	CULE	TYP	E: p	epti	.de			•				
	÷	(:	xi)	SEQU	ENCE	DES	CRIP	TION	: SE	Q II	) NO	23:	Ť			
50	Glu 1	Val	Gln	Leu	Gln 5		Ser	Gly	Val	Glu 1	_	Val	Arg	Ala	_	Ser 15

5	Ser	Val	Lys	Met 20	Ser	Cys	Lys	Ala	Ser 25	_	Tyr	Thr	Phé	Thr 3	Ser O	Asn
10	Gly	Ile	Asn 35	Trp	Val	Lys	Gln	Arg 40		Gly	Gln	Gly		Glu 5	Trp	Ile
	Gly	Tyr 50	Asn	Asn	Pro	Gly	Asn 55		Tyr	Ile	Ala		Asn O	Glu	Lys	Phe
15	Lys 65	Gly	Lys	Thr	Thr	Leu 70	Thr	Val	Asp	Lys	Ser 75	Ser	Ser	Thr	Ala	Tyr 80
20	Met	Gln	Leu	Arg	Ser 85	Leu	Thr	Ser	Glu	Asp 90	_	Ala	Val	Tyr		Cys 5
	Ala	Arg	Ser	Glu 100	Tyr	Tyr	Gly	Gly	Ser 10		Lys	Phe	Asp	Tyr 11		Gly
25	(24)		ORMA'							• •						
30			(i) :	~ () . ()	A) L B) T	ENGT YPE:	H: 1 ami		mino cid		ds					s. 1
		•	ii)   xi)				_	_		II Q	ON C	:24:				
35	Glu 1	Val	Gln	Leu	Val 5		Ser	Gly	Gly	Gly 1	-	Val	Gln	Pro	_	Arg 15
40	Ser	Leu	Arg	Leu 20		Cys	Ala	Ala	Ser 2		Phe	Thr	Phe	Asn 3	Asp 0	Tyr
	Ala	Met	His 35		Val	Arg	Gln	Ala 4		Gly	Lys	Gly		Glu 15	Trp	Val
45	Ser	Gly 50		Ser	Trp	Asp	Ser 5		Ser	Ile	Gly	_	Ala 50	Asp	Ser	Val
50	Lys 65	_	Arg	Phe	Thr	Ile 70	Ser	Arg	Asp	Asn	Ala 75		Asn	Ser	Leu	Ту <b>г</b> 80
	Leu	Gln	Met	Asn	Ser 85		Arg	Ala	Glu		Met 0	Ala	Leu	Tyr		Суз 95
55																

5	100 105 105 Tyr Asp Ser Gly Gly Tyr Phe Thr Val Ala
r	Phe Asp Ile Trp Gly 115
10	(25) INFORMATION FOR SEQ ID NO:25
15	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 111 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:25:
20	Asp Val Leu Met Thr Gln Thr Pro Leu Ser Leu Pro Val Ser Leu Gly 1 5 10 15
25	Asp Gln Ala Ser Ile Ser Cys Arg Ser Ser Gln Ile Ile Ile His Ser 20 25 30
30	Asp Gly Asn Thr Tyr Leu Glu Trp Phe Leu Gln Lys Pro Gly Gln Ser
	Pro Lys Leu Leu Ile Tyr Lys Val Ser Asn Arg Phe Ser Gly Val Pro 50 55 60
35	Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe Thr Leu Met Ile 65 70 75 80
	Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr Cys Phe Gln Gl 85 90 95
40	
	Ser His Val Pro His Thr Phe Gly Gly Gly Thr Lys Leu Glu Ile 100 105 110
45	(26) INFORMATION FOR SEQ ID NO:26
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 110 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:26:
55	

5	Glr 1	Ser	Val	Leu	Thr	Gln S	Pro	Pro	Ser	Ala 1	Ser .0.	Gly	Thr	Pro		Gln 15
10	Arg	val	Thr	Ile 20	Ser	Cys	Ser	Gly	Thr 2	Ser 5	Ser	Asn	Ile		Ser 30	Ser
	Thr	· Val	Asn 35	Trp	Туг	Gln	Gln	Leu 40	Pro	Gly	Met	Ala		Lys 15	Leu	Leu
15	Ile	Tyr 50	Arg	Asp	Ala	Met	Arg 55	Pro	Ser	Gly	Val	Pro 6	Asp 0	Arg	Phe	Ser
20						, 0					75					80
	Ser	Glu	Asp	Glu	Thr 85	Asp	Tyr	Tyr	Cys	Ala 90	Ala O	Trp	Ąsp	Val		Leu 95
25	Asn	Ala	Tyr	Val 100	Phe	Gly	Thr	Gly	Thr 105	Lys	Va]	Thr	· Vai	l Leu 110		
	(27)	INF	ORMA!	TION	FOR	SEQ	ID	NO:2	7							
30			(i) :	( E	1) Li 3) Ti	ENGT YPE:	RACT H: 1: ami: OGY:	ll a no a	mino cid	: aci	ds					. •
35	'a .	•		MOLEC SEQUE			_			Q ID	NO:	27:				
40	Gln 1	Val									Leu		Val	Thr		Gly .5
	Gln	Gln	Ala	Ser 20	Ile	Ser	Cys	Arg	Ser 25	Ser	Gln	Ile	Ile	Ile 3		Ser
45	Asp	Gly	Asn 35	Thr '	Tyr	Leu	Glu	Trp 40	Phe	Leu	Gln	Lys	Pro 4		Gln	Ser
50	Pro	<b>Lys</b> 50	Leu	Leu :	Ile <sub>.</sub>	Tyr	<b>Lys</b> 55	Val	Ser	Asn	Arg	Phe 60	_	Gly	Val	Pro
	Asp 65	Arg	Phe	Ser	Gly	S <b>er</b> 70	Gly .	Ser	Gly	Thr	Ser 75	Phe	Thr	Leu .	Ala	Ile 80
55																

5	s r	Arg	Val	Glu	Ala 85	Glu	Asp	Glu	Gly	Val ' 90		Tyr	Cys	Phe	Gln 9	
	Ser	His	Val	Pro 100	His	Thr	Phe	Gly	Gly 105	Gly	Thr	Lys	Leu	Glu 110		•
10	(28)	INFO	ORMA:	rion	FOR	SEQ	ID	NO: 2	8							
15		ı	(i) :	~ (2 (1	B) T	ENGT YPE:	H: 1 ami		mino cid	: aci	ds					
		(:	ii) 1	MOLE	CULE	TYP	E: p	epti	de	-						
		(:	xi)	SEQU	ENCE	DES	CRIF	TION	: SE	Q ID	NO:	28:				
20	Asp 1	Val	Val	Met	Thr 5		Ser	Pro	Leu	Ser 10		Pro	Val	Thr		Gly 5
25	Gln	Pro	Ala	Ser 20		Ser	Cys	Arg	Ser 25	Ser 5	Gln	Ser	Leu		Tyr 0	Ser
	Asp	Gly	Asn 35		Tyr	Leu	Asn	Trp		Gln	Gln	Arg		Gly 5	Gln	Ser
30	Pro	Arg 50		Leu	Ile	Tyr	Lys 5		Ser	Asn	Arg		Ser 0	Gly	Val	Pro
35	Asp 65	Arg	Phe	Ser	Gly	Ser 70		Ser	Glý	Thr	Asp 75	Phe	Thr	Leu	Lys	Ile 80
	Ser	Arg	Val	Glu	Ala 85		qaA	Val	Gly	Val		Tyr	Cys	Met	Gln	Gly 95
40	Thr	His	_Trp	Ser 100		Thr	· Phe	Gly	Gln 10	Gly 5	Thr	Lys	Val	Glu 1	Ile 10	Lys
45	(29)	INF	ORMA	TION	FOF	R SE	Q ID	ио:	29							
			(i)		JENCI (A) I (B) I (C) I	LABE PENG	TH: : am	111 . ino .	amina acid	o ac	ids					
50		(	(ii)	MOL	ECULI	E TY	PE:	pept	ide							
			(xi)	SEQ	UENC	E DE	SCRI	PTIO	พ: ร	EQ I	D NO	:29:				•

**53** 

5	Asp 1	Val	Leu	Met	Thr 5		Ser	Pro	Leu	Ser 1		Pro	Val	Thr		Gly .5
10	Gln	Pro	Ala	Ser 20	Ile	Ser	Cys	Arg	Ser 25		Gln	Ile	Ile		His O	Ser
	Asp	Gly	Asn 35	Thr	Tyr	Leu	Glu	Trp 40		Gln	Gln	Arg		Gly 5	Gln	Ser
15	Pro	Arg 50	Leu	Leu	Ile	Tyr	Lys 55		Ser	Asn	Arg		Ser O	Gly	Val	Pro
20	Asp 65	Arg	Phe	Ser	Gly	Ser 70	Gly	Ser	Gly	Thr	Asp 75	Phe	Thr	Leu	Lys	Ile 80
	Ser	Arg	Val	Glu	Ala 85	Glu	Asp	Val	Gly	Val 9	_	Tyr	Cys	Phe	_	Gly 5
25	Ser	His	Val	Pro 100	His	Thr	Phe	Gly	Gly 105	_	y Thi	Ly	s Va	1 Gl		<b>e</b> .
	(30)	INF	ORMA	TION	FOR	SEQ	ID	NO: 3	0							
30			(i)	Ċ	A) L B) T	ENGT YPE:	H: 1	.12 a .no a	mind		ids					
35		•		MOLE SEQU			-	•		EQ II	ON O	:30:				
	Asp 1	Ile	Val	Met	Thr 5	_	Ser	Pro	Asp	_	Leu .0	Alá	Val	Ser		Gly 15
40	Glu	Arg	_Ala_	Thr 20		Asn	Cys	Lys	Ser 2		Gln	Ser	Val	_	Tyr 30	Ser
45	Ser	Asn	Asn 35	Lys	Asn	Tyr	Leu	Ala 4	_	Tyr	Gln	Gln		Pro 15	Gly	Gln
50	Pro	Pro 50	_	Leu	Leu	Ile	Tyr 5		Ala	Ser	Thr		Glu 50	Ser	Gly	Val

5	Ile Ser Ser Leu Gln Ala Glu Asp Val Ala Val Tyr 3 85 90	Tyr Cys Gln Gln 95
10	Tyr Asp Thr Ile Pro Thr Phe Gly Gly Gly Thr Lys V	Val Glu Ile Lys 110
	(31) INFORMATION FOR SEQ ID NO:31	
15	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 111 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: peptide	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:31:	
	Asp Val Leu Met Thr Gln Thr Pro Asp Ser Leu Pro V 1 5 10	Val Ser Leu Gly 15
25	Asp Arg Ala Ser Ile Ser Cys Arg Ser Ser Gln Ile 1 20 25	Ile Ile His Ser 30
30	Asp Gly Asn Thr Tyr Leu Glu Trp Phe Leu Gln Lys I 35 40	Pro Gly Gln Ser 45
	Pro Lys Leu Leu Ile Tyr Lys Val Ser Asn Arg Phe S 50 55 60	
35	Asp Arg Phe Ser Gly Ser Gly Ser Gly Thr Asp Phe 1 65 70 75	Thr Leu Met Ile 80
40	Ser Arg Val Glu Ala Glu Asp Leu Gly Val Tyr Tyr ( 85 90	Cys Phe Gln Gly 95
	Ser His Val Pro His Thr Phe Gly Gly Thr Lys	Leu Glu Ile 110
45	(32) INFORMATION FOR SEQ ID NO:32	
50	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 117 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear	
<b>50</b>	(ii) MOLECULE TYPE: peptide	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:32:	
55		

1		GIII	rea	V <b>a</b> 1	GIU	ser	GIĀ	GIĄ	Gly 1	Leu 0	Val	Gln	Pro		Gİy 15
Ser	Arg	Lys	Leu 20	Ser	Cys	Ala	Ala	Ser 2	Gly 5	Phe	Thr	Phe			Phe
Gly	Met	His 35	Trp	Val	Arg	Gln	Ala 40	Pro	Glu	Lys	Gly			Trp	Val
Ala	Tyr 50	Ile	Ser	Ser	Gly	Ser -55	Phe	Thr	Ile	Tyr	His 6	Ala O	Asp	Thr	Val
•					70					75					80
Leu	Gln	Met	Thr	Ser 85	Leu	Arg	Ser	Glu	Asp 90	Thr	Ala	Met	Tyr		Cys 5
Ala	Arg	Met	Arg 100	Lys	Gly	Tyr	Ala	Met 105	Asp	Tyr	Trp	Gly			Thr
Thr		115													د
(33)			SEQUE ( <i>P</i> (E	ENCE () Li () T	CHAI ENGTI (PE:	RACT H: 1 ami	ERIS 25 a: no a	TICS mino cid		ds					
•	•				-						·				
Glu 1									Gly	Val		Gln	Pro		Arg 5
Ser	Leu	Arg	Leu 20	Ser	Cys	Ser	Ser	Ser 25	Gly	Phe	Ile	Phe			Tyr
Ala	Met '	Tyr 35	Trp '	Val .	Arg	Gln	Ala 40	Pro	Gly	Lys	Gly			Trp	Val
															Val
	Gly Ala Lys 65 Leu Ala Thr (33)	Ser Arg Gly Met Ala Tyr 50 Lys Gly 65 Leu Gln Ala Arg Thr Val (33) INFO (in) (in) (in) (in) Ser Leu Ser Leu	Ser Arg Lys  Gly Met His 35  Ala Tyr Ile 50  Lys Gly Arg 65  Leu Gln Met  Ala Arg Met  Thr Val Thr 115  (33) INFORMAT  (i) S  (ii) M  (xi) S  Glu Val Gln 1  Ser Leu Arg	Ser Arg Lys Leu 20  Gly Met His Trp 35  Ala Tyr Ile Ser 50  Lys Gly Arg Phe 65  Leu Gln Met Thr  Ala Arg Met Arg 100  Thr Val Thr Val 115  (33) INFORMATION  (i) SEQUE (A) (A) (A) SEQUE (	Ser Arg Lys Leu Ser 20  Gly Met His Trp Val 35  Ala Tyr Ile Ser Ser 50  Lys Gly Arg Phe Thr 65  Leu Gln Met Thr Ser 85  Ala Arg Met Arg Lys 100  Thr Val Thr Val Ser 115  (33) INFORMATION FOR  (i) SEQUENCE (A) Li (B) Ti (C) To  (ii) MOLECULE (xi) SEQUENCE Glu Val Gln Leu Val 1  Ser Leu Arg Leu Ser 20  Ala Met Tyr Trp Val	Ser Arg Lys Leu Ser Cys 20  Gly Met His Trp Val Arg 35  Ala Tyr Ile Ser Ser Gly 50  Lys Gly Arg Phe Thr Ile 65 70  Leu Gln Met Thr Ser Leu 85  Ala Arg Met Arg Lys Gly 100  Thr Val Thr Val Ser 115  (33) INFORMATION FOR SEQ (i) SEQUENCE CHAM (A) LENGTM (B) TYPE: (C) TOPOLO (ii) MOLECULE TYPE (xi) SEQUENCE DESC Glu Val Gln Leu Val Gln 1  Ser Leu Arg Leu Ser Cys 20  Ala Met Tyr Trp Val Arg	Ser Arg Lys Leu Ser Cys Ala 20  Gly Met His Trp Val Arg Gln 35  Ala Tyr Ile Ser Ser Gly Ser 50  Lys Gly Arg Phe Thr Ile Ser 65  Ala Arg Met Thr Ser Leu Arg 85  Ala Arg Met Arg Lys Gly Tyr 100  Thr Val Thr Val Ser 115  (33) INFORMATION FOR SEQ ID  (i) SEQUENCE CHARACT (A) LENGTH: 1 (B) TYPE: ami (C) TOPOLOGY:  (ii) MOLECULE TYPE: po (xi) SEQUENCE DESCRIP  Glu Val Gln Leu Val Gln Ser 1  Ser Leu Arg Leu Ser Cys Ser 20  Ala Met Tyr Trp Val Arg Gln	Ser Arg Lys Leu Ser Cys Ala Ala 20  Gly Met His Trp Val Arg Gln Ala 35  Ala Tyr Ile Ser Ser Gly Ser Phe 50  Lys Gly Arg Phe Thr Ile Ser Arg 65  Leu Gln Met Thr Ser Leu Arg Ser 85  Ala Arg Met Arg Lys Gly Tyr Ala 100  Thr Val Thr Val Ser 115  (33) INFORMATION FOR SEQ ID NO:3  (i) SEQUENCE CHARACTERIS (A) LENGTH: 125 a. (B) TYPE: amino a (C) TOPOLOGY: lin  (ii) MOLECULE TYPE: peptic (xi) SEQUENCE DESCRIPTION  Glu Val Gln Leu Val Gln Ser Gly 1  Ser Leu Arg Leu Ser Cys Ser Ser 20  Ala Met Tyr Trp Val Arg Gln Ala	Ser Arg Lys Leu Ser Cys Ala Ala Ser  20  Gly Met His Trp Val Arg Gln Ala Pro 35  Ala Tyr Ile Ser Ser Gly Ser Phe Thr 50  Lys Gly Arg Phe Thr Ile Ser Arg Asp 65  Leu Gln Met Thr Ser Leu Arg Ser Glu 85  Ala Arg Met Arg Lys Gly Tyr Ala Met 100  Thr Val Thr Val Ser 115  (33) INFORMATION FOR SEQ ID NO:33  (i) SEQUENCE CHARACTERISTICS (A) LENGTH: 125 amino (B) TYPE: amino acid (C) TOPOLOGY: linear  (ii) MOLECULE TYPE: peptide  (xi) SEQUENCE DESCRIPTION: SEC  Glu Val Gln Leu Val Gln Ser Gly Gly 1  Ser Leu Arg Leu Ser Cys Ser Ser Ser 20  Ala Met Tyr Trp Val Arg Gln Ala Pro	Ser Arg Lys Leu Ser Cys Ala Ala Ser Gly 20  Gly Met His Trp Val Arg Gln Ala Pro Glu 35  Ala Tyr Ile Ser Ser Gly Ser Phe Thr Ile 50  Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn 65  Leu Gln Met Thr Ser Leu Arg Ser Glu Asp 85  Ala Arg Met Arg Lys Gly Tyr Ala Met Asp 100  Thr Val Thr Val Ser 115  (33) INFORMATION FOR SEQ ID NO:33  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 125 amino aci (B) TYPE: amino acid (C) TOPOLOGY: linear  (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID  Glu Val Gln Leu Val Gln Ser Gly Gly Gly 1  Ser Leu Arg Leu Ser Cys Ser Ser Ser Gly 20  Ala Met Tyr Trp Val Arg Gln Ala Pro Gly	Ser Arg Lys Leu Ser Cys Ala Ala Ser Gly Phe  20  Gly Met His Trp Val Arg Gln Ala Pro Glu Lys  35  Ala Tyr Ile Ser Ser Gly Ser Phe Thr Ile Tyr  50  Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Pro 65  70  Ala Arg Met Thr Ser Leu Arg Ser Glu Asp Thr 85  Ala Arg Met Arg Lys Gly Tyr Ala Met Asp Tyr 100  Thr Val Thr Val Ser 115  (33) INFORMATION FOR SEQ ID NO:33  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 125 amino acids (B) Type: amino acids (C) TOPOLOGY: linear  (ii) MOLECULE TYPE: peptide  (xi) SEQUENCE DESCRIPTION: SEQ ID NO:  Glu Val Gln Leu Val Gln Ser Gly Gly Gly Val  1  Ser Leu Arg Leu Ser Cys Ser Ser Ser Gly Phe 20  Ala Met Tyr Trp Val Arg Gln Ala Pro Gly Lys	Ser Arg Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr  20  Gly Met His Trp Val Arg Gln Ala Pro Glu Lys Gly  Ala Tyr Ile Ser Ser Gly Ser Phe Thr Ile Tyr His  50  Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Pro Lys  65  Leu Gln Met Thr Ser Leu Arg Ser Glu Asp Thr Ala  85  Ala Arg Met Arg Lys Gly Tyr Ala Met Asp Tyr Trp  100  Thr Val Thr Val Ser  115  (33) INFORMATION FOR SEQ ID NO:33  (i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 125 amino acids (B) Type: amino acids (C) TOPOLOGY: linear  (ii) MOLECULE TYPE: peptide  (xi) SEQUENCE DESCRIPTION: SEQ ID NO:33:  Glu Val Gln Leu Val Gln Ser Gly Gly Gly Val Val  1  Ser Leu Arg Leu Ser Cys Ser Ser Ser Gly Phe Ile  20  Ala Met Tyr Trp Val Arg Gln Ala Pro Gly Lys Gly	Ser Arg Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe  20  Gly Met His Trp Val Arg Gln Ala Pro Glu Lys Gly Leu  35  Ala Tyr Ile Ser Ser Gly Ser Phe Thr Ile Tyr His Ala 50  Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Pro Lys Asn 65  70  Leu Gln Met Thr Ser Leu Arg Ser Glu Asp Thr Ala Met 85  Ala Arg Met Arg Lys Gly Tyr Ala Met Asp Tyr Trp Gly 100  Thr Val Thr Val Ser 115  (33) INFORMATION FOR SEQ ID NO:33  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 125 amino acids (B) Type: amino acids (C) TOPOLOGY: linear  (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:33:  Glu Val Gln Leu Val Gln Ser Gly Gly Gly Val Val Gln 1  Ser Leu Arg Leu Ser Cys Ser Ser Ser Gly Phe Ile Phe 20  Ala Met Tyr Trp Val Arg Gln Ala Pro Gly Lys Gly Leu	Ser Arg Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser 20 25 3  Gly Met His Trp Val Arg Gln Ala Pro Glu Lys Gly Leu Glu 45 40 45  Ala Tyr Ile Ser Ser Gly Ser Phe Thr Ile Tyr His Ala Asp 50 55 60  Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Pro Lys Asn Thr 75  Leu Gln Met Thr Ser Leu Arg Ser Glu Asp Thr Ala Met Tyr 85 90  Ala Arg Met Arg Lys Gly Tyr Ala Met Asp Tyr Trp Gly Gln 100 105 11  Thr Val Thr Val Ser 115  (33) INFORMATION FOR SEQ ID NO:33  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 125 amino acids (B) TypE: amino acid (C) TOPOLOGY: linear  (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:33:  Glu Val Gln Leu Val Gln Ser Gly Gly Gly Val Val Gln Pro 10  Ser Leu Arg Leu Ser Cys Ser Ser Ser Gly Phe Ile Phe Ser 20 25 36  Ala Met Tyr Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Ser Cys Trp Cala Arg Cln Ala Pro Gly Lys Gly Leu Glu Ser Cys Trp Cala Arg Cln Ala Pro Gly Lys Gly Leu Glu Ser Cys Trp Cala Arg Cln Ala Pro Gly Lys Gly Leu Glu Ser Cys Trp Cala Arg Cln Ala Pro Gly Lys Gly Leu Glu Ser Cys Cala Cala Cala Cala Cala Cala Cala Cal	Ser Arg Lys Leu Ser Cys Ala Ala Ser Gly Phe Thr Phe Ser Ser 20  Gly Met His Trp Val Arg Gln Ala Pro Glu Lys Gly Leu Glu Trp 40  Ala Tyr Ile Ser Ser Gly Ser Phe Thr Ile Tyr His Ala Asp Thr 50  Lys Gly Arg Phe Thr Ile Ser Arg Asp Asn Pro Lys Asn Thr Leu 65  To 70  Leu Gln Met Thr Ser Leu Arg Ser Glu Asp Thr Ala Met Tyr Tyr 85  Ala Arg Met Arg Lys Gly Tyr Ala Met Asp Tyr Trp Gly Gln Gly 100  Thr Val Thr Val Ser 115  (33) INFORMATION FOR SEQ ID NO:33  (i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 125 amino acids (B) Type: amino acid (C) TOPOLOGY: linear  (ii) MOLECULE Type: peptide  (xi) SEQUENCE DESCRIPTION: SEQ ID NO:33:  Glu Val Gln Leu Val Gln Ser Gly Gly Gly Val Val Gln Pro Gly 1  Ser Leu Arg Leu Ser Cys Ser Ser Ser Gly Phe Ile Phe Ser Ser 20  Ala Met Tyr Trp Val Arg Gln Ala Pro Gly Lys Gly Leu Glu Trp

5	Lys 65	Gly	Arg	Phe	Thr	Ile 70	Ser	Arg	Asn	Asp	Ser 75	Lys	Asn	Thr	Leu	Phe 80
•	Leu	Gln	Met	Asp	Ser 85	Leu	Arg	Pro	Glu	Asp 90		Gly	Val	Tyr	Phe 9	Cys 5
10	Ala	Arg	Asp	Gly 100	Gly	His	Gly	Phe	Cys 105		Ser	Ala	Ser	Cys 11	Phe 0	Gly
15		Asp	115	_				120		Val	Thr	Va]	Sei 125			
	(34)	INFO	ORMA:	rion	FOR	SEQ	ID	NO: 3	4							
20			(i) :	()	A) L	ENGT YPE:	H: 1 ami	.17 a	mino cid		.ds					
		(	ii) I	MOLE	CULE	TYP	E: p	epti	.de							
25		(:	xi)	SEQU	ENCE	DES	CRIF	TION	: SE	Q II	NO:	34:				
	Glu 1	Val	Gln	Leu	Val 5		Ser	Gly	Gly	Gly 1		Val	Gln	Pro	Gly 1	Arg .5
30	Ser	Leu ·	Arg	Leu 20		Cys	Ala	Ala	Ser 2		Phe	Ile	Phe		Ser 80	Phe
35	Gly	Met	His 35		Val	Arg	Gln	Ala 4		Gly	Lys	Gly		Glu 15	Trp	Val
	Ala	Tyr 50		Ser	Ser	Asp	Gly 5		Thr	Ile	Tyr	His 6	Ala 50	Asp	Ser	Val
<b>1</b> 0	Lys 65		_Arq	Phe	Thr	Ile 70		Arg	Asp	Asp	Pro 75		Asn	Thr	Leu	Phe 80
15	Leu	Gln	Met	Thr	Ser 85		. Arg	Ser	Glu	Asp 9	Thr O	Ala	Met	Tyr	Tyr	Cys 95
	Ala	Arg	Met	Arg		Gly	Tyr	Ala	Met 10	Asp 5	Tyr	Trp	Gly	Gln 1	Gly 10	Thr
50	Thr	· Val	Thr 115		l Sei	r										
	(35)	INE	FORM	TIO	4 FO	R SE	Q ID	ио:	35							
_																

5			(+)	(	A) L B) T	ENGT YPE:	H: 1 ami	.20 a	mino		ids					
		(	ii)	MOLE	CULE	TYP	E: p	epti	.de							
10		(	xi)	SEQU	ENCE	DES	CRIP	MOIT	: SE	Q II	NO:	35:				
	Gln 1	Val	Gln	Leu	Val 5	Glu	Ser	Gly	Gly	Gly 1		Val	Gln	Pro		Arg L5
15	/Ser	Leu	Arg	Leu 20	Ser	Cys	Ala	Ala	Ser 29		Phe	Thr	Phe	_	Ser 0	Туг
20	Ala	Met	His 35	Trp	Val	Arg	Gln	Ala 40		Gly	Lys	Gly		Glu 5	Trp	Val
25	Ala	Val 50	Ile	Ser	Tyr	Asp	Gly 55		Asn	Lys	Tyr		Ala O	Asp	Ser	Val
	Lys 65	Ġly	Arg	Phe	Thr	Ile 70	Ser	Arg	Asp	Asn	Ser 75	Lys	Asn	Thr	Leu	Tyr 80
30	Leu	Gln	Met	Asn	Ser 85	Leu	Arg	Ala	Glu	Asp 90	_	Ala	Val	Tyr		Cys 5
35	Ala	Arg	Asp	Arg 100	Lys	Asp	Trp	Gly	Trp 10		Leu	Phe	Asp	Tyr 11		Gly
	Gln	Gly	Thr 115	Leu	Val	Thr	Val	Ser 120								
40	(36)	INF	ORMA	TION	FOR	SEQ	ID	<b>NO:</b> 3	6							
			(i)_	(I	A) L B) T	engt YPE:	RACT H: 1 ami OGY:	17 a	mino		lds					
45		(	ii)	MOLE	CULE	TYP	E: p	epti	.de							
		(:	xi)	SEQU	ENCE	DES	CRIP	TION	: SE	EQ II	NO:	36:	5			
50	Gln 1	Val	Gln	Leu	Val 5		Ser	Gly	Gly	Gly 1	_	Val	Gln	Pro		Arg 15

5	Ser	Leu	Arg	Leu 20	Ser	Cys	Ala	Ala	Ser 2	Gly 5	Phe	Thr	Phe		Ser 10	Phe
	Gly	Met	His 35	Trp	Val	Arg	Gln	Ala 40	Pro )	Gly	Lys	Gly		Glu 5	Trp	Val
10	Ala	Tyr 50	Ile	Ser	Ser	Gly	Ser 55	Phe	Thr	Ile	Tyr		Ala O	Asp	Ser	Val
15	Lys 65	Gly	Arg	Phe	Thr	Ile 70	Ser	Arg	Asp	Asn	Ser 75	Lys	Asn	Thr	Leu	Tyr 80
20	Leu	Gln	Met	Asn	Ser 85	Leu	Arg	Ala	Glu	Asp 90		Ala	Val	Tyr		Cys 5
20	Ala	Arg	Met	Arg 100	Lys	Gly	Tyr	Ala	Met 105		Tyr	Trp	Gly	Gln 11		Thr
25	Leu	Val	Thr 115	Val	Ser				٠				•			
	(37)	INF	ORMA	TION	FOR	SEQ	ID	NO: 3	7							
30			(i) :	(1	A) Li B) Ti	ENGT:	H: 9 amai		ino cid	: acid	ls					
		(:	ii) I	MOLE	CULE	TYP	E: p	epti	de							
35		(2	ki) :	SEQUI	ENCE	DES	CRIP	TION	: SE	Q ID	NO:	37:				
	Glu 1	Val	Gln	Leu	Va1 5	Glu	Ser	Gly	Gly	Gly 1		Val	Gln	Pro		Gly .5
<b>4</b> 0	Ser	Leu	Arg	Leu 20	Ser	Cys	Ala	Ala	Ser 25	_	Phe	Thr	Phe	_	Ser 0	Tyr
45	Trp	Met	Ser 35	Trp	Val	Arg	Gln	Ala 40		Gly	Lys	Gly	Leu 4	_	Trp	Val
50	Åla	Asn 50	Ile	Lys	Gln	Asp	Gly 55		Glu	Lys	Tyr	Tyr 6	_	Asp	Ser	Val
	Lys 65	Gly	Arg	Phe	Thr	Ile 70	Ser	Arg	Asp	Asn	Ala	Lys	Asn	Ser	Leu	Tyr 80

_	Ded	GIII	mec	ASI	85	Leu	Arg	АТА	GIU	Asp 90		Ата	Vai	Tyr	_	Cys 95
5	Ala	Arg														
10	(38)	INFO	ORMA'	rion	FOR	SEQ	ID	мо: 3	8							
45		ı	(i) £	(1	A) L B) T	ENGT YPE:	RACT H: 1 ami OGY:	17 a no a	minc		.ds					
15		(:	ii) 1	MOLE	CULE	TYP	E: p	epti	de							
.£.		(2	ki) s	SEQUI	ENCE	DES	CRIP	TION	: SE	Q II	NO:	38:				
20	Glu 1	Val	Gln	Leu	Val 5		Ser	Gly	Gly	Gly 1		Val	Gln	Pro		Gly L5
. 25	Ser	Leu	Arg	Leu 20	Ser	Cys	Ala	Ala	Ser 2		Phe	Thr	Phe		Ser 0	Phe
	Gly	Met	His 35	Trp	Val	Arg	Gln	Ala 40		Gly	Lys	Gly		Glu 5	Trp	Val
30	Ala	Tyr 50	Ile	Ser	Ser	Gly	Ser 55		Thr	Ile	Tyr	_	Ala O	Asp	Ser	Val
35	Lys 65	Gly	Arg	Phe	Thr	Ile 70	Ser	Arg	Asp	Asn	Ala 75	Lys	Asn	Thr	Leu	Phe 80
	Leu	Gln	Met	Thr	Ser 85		Arg	Ala	Glu	Asp 9		Ala	Met	Tyr	_	Cys 95
40	Ala	Arg.	_Met_	Arg 100	Lys	Gly	Tyr	Ala	Met 10		Tyr	Trp	Gly		Gly LO	Thr
45	Thr	Val	Thr 115		Ser	· .						÷				
	(39)	INF	ORMA	TION	FOR	SEC	D ID	NO:3	39							
50			(i)	(	A) L B) T	ENGT	RACT TH: 1 ami	l5 ar ino a	nino acid	aci	ds					
		(	ii)	MOLE	CULE	TYI	PE: I	ept:	ide							

		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:39:
5	Met 1	Gly Trp Ser Cys Ile Ile Leu Phe Leu Val Ala Thr Ala Thr 5 10 15
	(40)	INFORMATION FOR SEQ ID NO:40
10		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:40:
20	Lys 1	Thr Ser Leu Arg Pro Gly Lys Gly Ser Ser Asp Tyr Glu Lys Lys 5 10 15
	(41)	INFORMATION FOR SEQ ID NO:41
25		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
30		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:41:
	Lys 1	Thr Ser Leu Arg Pro Gly Lys Gly Ser Ser Glu Tyr Glu Lys Lys 5 10 15
35	(42)	INFORMATION FOR SEQ ID NO:42
<b>4</b> 0		(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
		$(\bar{1}i)$ MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:42:
45	Gln 1	Thr Ser Leu Arg Pro Asp Lys Gly Ser Ser Asp His Glu Lys Lys 5 10 15
50	(43)	INFORMATION FOR SEQ ID NO:43
		(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
55		

5	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:43:
10	Gln Thr Ser Leu Arg Pro Asp Lys Gly Ser Ser Asp Gln Glu Lys Lys 1 5 10 15
	(44) INFORMATION FOR SEQ ID NO:44
15	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:44:
	Gln Ser Ser Leu Arg Pro Asp Lys Gly Ser Ser Asp Gln Glu Lys Lys 1 5 10 15
25	(45) INFORMATION FOR SEQ ID NO:45
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:45:
35	Gln Thr Ser Leu Arg Pro Asp Lys Gly Ser Ser Asp Pro Glu Lys Lys  1 5 10 15
	(46) INFORMATION FOR SEQ ID NO:46
40	(i) SEQUENCE CHARACTERISTICS:  _ (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:46:
50	Gln Thr Ser Leu Arg Pro Asp Lys Gly Ser Ser Asp Pro Glx Lys Lys 1 5 10 15
	(47) INFORMATION FOR SEQ ID NO:47
	(i) SEQUENCE CHARACTERISTICS:
55	

5		<ul><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>	
		(ii) MOLECULE TYPE: peptide	
10		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:47:	
,,	Gln 1	Thr Ser Leu Arg Pro Asp Lys Gly Ser Ser Asp Pro Glu Lys Th 5 10 15	r
15	(48)	INFORMATION FOR SEQ ID NO:48	
		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>	
20		(ii) MOLECULE TYPE: peptide	
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:48:	
25	Gln 1	Thr Ser Leu Arg Ala Asp Lys Gly Ser Ser Asp Gln Glu Lys Ly 5 10 15	s
	(49)	INFORMATION FOR SEQ ID NO:49	
30		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>	
35		(ii) MOLECULE TYPE: peptide	
55		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:49:	
	Gln 1	Thr Ser Leu Arg Pro Asp Lys Gly Lys Ser Asp Ser Glu Lys Ly 5 10 15	s
40	(50)	INFORMÁTION FOR SEQ ID NO:50	
45		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>	
		(ii) MOLECULE TYPE: peptide	
50		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:50:	
	Gln 1	Thr Ser Leu Arg Pro Ala Arg Gly Ser Ser Asp Gln Glu Lys Ly 5 10 15	rs

5	(51)	INFORMATION FOR SEQ ID NO:51
		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
10		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:51:
15	Gln 1	Thr Ser Leu Lys Pro Gly Arg Gly Ser Ser Asp Pro Glu Lys Lys 5 10 15
	(52)	INFORMATION FOR SEQ ID NO:52
20		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
25		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:52:
	Gln 1	Thr Ser Leu Arg Pro Gly Arg Gly Ser Ser Asp Thr Glu Lys Lys 5 10 15
30	(53)	INFORMATION FOR SEQ ID NO:53
35		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	ner Spatial is	(ii) MOLECULE TYPE: peptide
40		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:53:
40	Gln 1	Ile_Ser_Leu Arg Pro Gly Lys Gly Ser Ser Asp Ser Glu Lys Lys 5 10 15
45	(54)	INFORMATION FOR SEQ ID NO:54
	·	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:54:

5	Gln 1	Thr Se	r Leu <i>l</i>	Arg I	Pro Gly	y Lys	Gly	Asp 10		Asp	Glu	Asp	Lys Lys 15
	(55)	INFORM	ATION	FOR	SEQ ID	NO:5	5						
10		(i)	(A . (B	) LE ) TY	CHARAC NGTH: PE: am POLOGY	16 am ino a	ino cid		S				
		(ii)	MOLEC	ULE	TYPE:	pepti	de		٠				
15		(xi)	SEQUE	NCE	DESCRI	PTION	: SE	Q ID	NO:	55:			
	Glu 1	Thr Al	a Leu <i>l</i>	Arg I 5	Pro Gly	y Lys	Gly	Ala 10		Asp	Ala	Asp	Lys Lys 15
20	(56)	INFORM	ATION	FOR	SEQ ID	NO:5	6						
25		(i)	•	) LE ) TY	CHARAC NGTH: PE: am POLOGY	16 am	ino	-	s				
		(ii)	MOLEC	ULE	TYPE:	pepti	de						
30	Val 1		SEQUE a Leu <i>l</i>						Ser		Glu	Asp	Lys Lys 15
	_			Ū									
35	(57)	INFORM											
		(i)	•	) LE	CHARAC INGTH: IPE: an IPOLOGY	16 am	ino cid		ls				
40		(ii)	_MOLEC	ULE	TYPE:	pepti	.de						
		(xi)	SEQUE	NCE	DESCRI	PTION	: SE	Q ID	NO:	:57:			
45	Val 1	Thr Al	a Leu i	Arg 1	Pro Gl	y Lys	Gly	Ala 1		Asp	Glu	Glu	Lys Lys 15
	(58)	INFORM	MOITA	FOR	SEQ II	NO: 5	8						
50		(i)	(B	) LE	CHARAC EMGTH: (PE: an OPOLOG)	16 am	nino ncid		ls				
55													

		(ii) MOLECULE TYPE: peptide
5		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:58:
10	Val 1	Thr Ala Leu Arg Pro Gly Lys Gly Ala Ser Asx Ala Asx Lys Lys 5 10 15
10	(59)	INFORMATION FOR SEQ ID NO:59
15	• ••	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
		(ii) MOLECULE TYPE: peptide
20		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:59:
	Val 1	Thr Ala Leu Arg Pro Gly Lys Gly Ala Ser Asp Glu Asp Asp Glu 5 10 15
25	(60)	INFORMATION FOR SEQ ID NO:60
30		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:60:
35	Gln 1	Thr Ser Leu Arg Pro Asp Lys Gly Ser Ser Asp Gln Glu Thr Thr 5 10 15
	. (61)	INFORMATION FOR SEQ ID NO:61
40		(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:61:
50	Gln 1	Asn Ser Leu Thr Pro Gly Lys Gly Ser Ser Ser Pro Glu Lys Lys 5 10 15
	(62)	INFORMATION FOR SEQ ID NO:62
		(i) SEQUENCE CHARACTERISTICS:
55		· ·

5		<ul><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:62:
10	Val 1	Thr Lys Val Arg Pro Gly Lys Gly Asp Ser Asp Ser Asp Lys Lys 5 10 15
15	(63)	INFORMATION FOR SEQ ID NO:63
		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
20		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:63:
25	Val 1	Thr Lys Val Arg Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
	(64)	INFORMATION FOR SEQ ID NO:64
30		(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
35		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:64:
40	Val 1	Thr Arg Val Arg Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
	(65)	INFORMATION FOR SEQ ID NO:65
45		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
50		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:65:
	Leu 1	Thr Lys Val Arg Pro Gly Lys Gly Asp Ser Asp Ser Glu Lys Lys 5
55		

	(66)	INFORMATION FOR SEQ ID NO:66
5		<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 16 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul>
10		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:66:
15	Val 1	Thr Lys Val Arg Pro Gly Lys Gly Asp Ser Asp Ser Glu Gln Lys 5 10 15
	(67)	INFORMATION FOR SEQ ID NO:67
20		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
25		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:67:
	Val 1	Thr Lys Val Arg Pro Glu Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
30	(68)	INFORMATION FOR SEQ ID NO:68
35		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
40		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:68:
••	Val 1	Thr_Lys_Val Arg Pro Glu Lys Gly Asp Ser Asp Ser Glu Lys Lys 5 10 15
45	(69)	INFORMATION FOR SEQ ID NO:69
		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:69:
55		

5	val 1	The Lys Val Ser Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys  10 15
	(70)	INFORMATION FOR SEQ ID NO:70
10		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:70:
20	Val 1	Thr Lys Val Arg Ser Gly Lys Gly Glu Ser Asp Ala Glu Lys Lys 5 10 15
	(71)	INFORMATION FOR SEQ ID NO:71
<i>2</i> 5		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
30		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:71:
	Val 1	Thr Ser Val Lys Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
35	(72)	INFORMATION FOR SEQ ID NO:72
40		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	•	(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:72:
<b>4</b> 5	Val	Ser Ser Val Lys Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
50	(73)	INFORMATION FOR SEQ ID NO:73
55		(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear

		(ii) MOLECULE TYPE: peptide
5		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:73:
	Val 1	Thr Ser Ala Lys Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
10	(74)	INFORMATION FOR SEQ ID NO:74
15		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
20		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:74:
20	Val 1	Ser Ser Ala Lys Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
25	(75)	INFORMATION FOR SEQ ID NO:75
		<ul> <li>(i) SEQUENCE CHARACTERISTICS:         <ul> <li>(A) LENGTH: 16 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul> </li> </ul>
30		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:75:
35	Val 1	Thr Ser Ala Arg Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
	(76)	INFORMATION FOR SEQ ID NO:76
40		<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>- (A) LENGTH: 16 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul>
45		(ii) MOLECULE TYPE: peptide
40		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:76:
50	Val 1	Ser Pro Ala Lys Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
	(77)	INFORMATION FOR SEQ ID NO:77
		(i) SEQUENCE CHARACTERISTICS:
55		

5		<ul><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
10		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:77:
	Val 1	Thr Lys Ala Arg Pro Gly Lys Gly Asp Ser Asp Val Glu Lys Asn 5 10 15
15	(78)	INFORMATION FOR SEQ ID NO:78
20		(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
20		(ii) MOLECULE TYPE: peptide
•		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:78:
25	Val 1	Thr Leu Ile Pro Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
	(79)	INFORMATION FOR SEQ ID NO:79
30		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
35		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:79:
40	Val 1	Thr Leu Leu Gln Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
	(80)	INFORMATION FOR SEQ ID NO:80
45		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
50		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:80:
	Val 1	Thr Leu Leu Gln Pro Gly Lys Gly Asp Ser Asp Ala Asp Lys Lys 5 10 15

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5	(81)	INFORMATION FOR SEQ ID NO:81
		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
10		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:81:
15	Val 1	Thr Leu Leu Gln Pro Gly Lys Gly Asp Ser Asp Ala Glu Arg Lys 5 10 15
	(82)	INFORMATION FOR SEQ ID NO:82
20		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
25		(ii) MOLECULE TYPE: peptide
25		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:82:
٠	Val 1	Thr Leu Leu Gln Ala Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys $5$ $10$ $15$
30	(83)	INFORMATION FOR SEQ ID NO:83
35		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	-	(ii) MOLECULE TYPE: peptide
40		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:83:
	Val 1	Thr_Leu Leu Gln Pro Gly Glu Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
45	(84)	INFORMATION FOR SEQ ID NO:84
		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:84:

5	Leu 1	Thr Leu Leu Gln Pro Gly Asn Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
	(85)	INFORMATION FOR SEQ ID NO:85
10		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:85:
20	Val 1	Thr Leu Leu Gln Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Ile 5 10 15
	(86)	INFORMATION FOR SEQ ID NO:86
25		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
30		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:86:
	Val 1	Thr Leu Phe Gln Pro Gly Gln Gly Asp Ser Asp Pro Glu Lys Lys 5 10 15
35	(87)	INFORMATION FOR SEQ ID NO:87
40		<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 16 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul>
		( <u>i</u> i) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:87:
45	Val 1	Thr Leu Pro Gln Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 5 10 15
E0	(88)	INFORMATION FOR SEQ ID NO:88
50		<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 16 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul>
55		

5	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:88:
10	Val Thr Leu Pro Gln Pro Gly Lys Gly Asp Trp Asp Ala Glu Lys Lys 1 5 10 15
	(89) INFORMATION FOR SEQ ID NO:89
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:89:
	Val Thr Phe Leu Ser Pro Gly Gln Gly Asp Ser Asp Ala Glu Lys Lys 1 5 10 15
25	(90) INFORMATION FOR SEQ ID NO:90
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:90:
35	Glu Ser Ser Ala Arg Pro Gly Lys Gly Asp Ser Asp Ala Glu Lys Lys 1 5 10 15
40	(91) INFORMATION FOR SEQ ID NO:91
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:91:
50	Val Thr Leu Ser Ser Pro Gly Gln Gly Asp Ser Asp Ala Glu Lys Lys 1 5 10 15
	(92) INFORMATION FOR SEQ ID NO:92
55	(i) SEQUENCE CHARACTERISTICS:

5		<ul><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
10		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:92:
	Val 1	Thr Thr Ala Lys Pro Glu Lys Gly Asp Ser Asp Val Glu Lys Lys 5 10 15
15	(93)	INFORMATION FOR SEQ ID NO:93
20		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:93:
25	Val 1	Thr Thr Pro Lys Pro Asp Lys Gly Asp Ser Asp Val Glu Lys Lys 5 10 15
	(94)	INFORMATION FOR SEQ ID NO:94
30		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
35		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:94:
40	Val 1	Thr Ala Pro Arg Pro Gly Lys Gly Ala Ser Ser Ala Glu Lys Lys 5 10 15
	(95)	INFORMATION FOR SEQ ID NO:95
45		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
50		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:95:
	Val 1	Thr Ala Pro Lys Pro Gly Lys Gly Thr Ser Ser Ala Glu Lys Lys 5 10 15

	(96)	INFORMATION FOR SEQ ID NO:96
5		(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
10		(ii) MOLECULE TYPE: peptide
		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:96:
15	Val 1	Thr Thr Pro Lys Pro Gly Lys Gly Ala Ser Ser Ala Glu Lys Lys 5 10 15
	.∉(97)	INFORMATION FOR SEQ ID NO:97
20		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
		(ii) MOLECULE TYPE: peptide
25		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:97:
	Val 1	Ser Ala Pro Lys Pro Gly Lys Gly Ala Ser Ser Ala Glu Lys Lys 5 10 15
30	(98)	INFORMATION FOR SEQ ID NO:98
35		<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
-	•	(ii) MOLECULE TYPE: peptide
40		(xi) SEQUENCE DESCRIPTION: SEQ ID NO:98:
	Val	Thr_Ala Pro Arg Ser Gly Lys Gly Ala Ser Ser Ala Glu Lys Lys 5 10 15
45	(99)	INFORMATION FOR SEQ ID NO:99
	·	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50		(ii) MOLDON D. GUDD
		(ii) MOLECULE TYPE: peptide

5	Val Thr Ala Pro Lys Ser Gly Lys Gly Ala Ser Ser Ala Glu Lys Lys 1 5 10 15
	(100) INFORMATION FOR SEQ ID NO:100
10	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:100:
	Val Thr Ala Pro Lys Pro Asp Lys Gly Val Ser Ser Ala Glu Lys Lys 1 5 10 15
20	(101) INFORMATION FOR SEQ ID NO:101
25	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:101:
30	Val Thr Ala Pro Lys Ser Glu Lys Gly Val Ser Ser Ala Glu Lys Lys 1 5 10 15
35	(102) INFORMATION FOR SEQ ID NO:102
	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
40	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:102:
45	Phe Thr Ala Pro Lys Pro Gly Lys Gly Ala Ser Ser Ala Glu Lys Lys 1 5 10 15
	(103) INFORMATION FOR SEQ ID NO:103
50	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear

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	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:103:
	Leu Thr Ala Pro Lys Pro Gly Arg Gly Val Ser Ser Ala Glu Lys Lys 1 5 10 15
10	(104) INFORMATION FOR SEQ ID NO:104
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:104:
20	Val Thr Ala Pro Lys Ser Gly Lys Gly Ala Ser Ser Ala Glu Lys Arg 1 5 10 15
25	(105) INFORMATION FOR SEQ ID NO:105
	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
30	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:105:
35	Val Ser Ala Pro Lys Pro Gly Lys Glu Gly Ser Ser Ala Glu Lys Lys 1 5 10 15
	(106) INFORMATION FOR SEQ ID NO:106
40	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:106:
50	Val Thr Ala Pro Lys Pro Arg Lys Gly Ala Ser Ser Ala Glu Lys Lys 1 5 10 15
	(107) INFORMATION FOR SEQ ID NO:107
55	(i) SEQUENCE CHARACTERISTICS:

5	(A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:107:
10	Val Thr Phe Leu Ser Pro Gly Gln Gly Asn Ser Asp Ala Glu Leu Pro 1 5 10 15
15	(108) INFORMATION FOR SEQ ID NO:108
73	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:108:
25	Val Thr Phe Leu Ser Pro Gly Gln Gly Asn Ser Asp Glu Asp Leu Pro 1 5 10 15
	(109) INFORMATION FOR SEQ ID NO:109
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:109:
40	Val Thr Leu Ser Ser Pro Gln Arg Gly Asp Ser Asp Ala Glu Lys Lys 1 5 10 15
	(110) INFORMATION FOR SEQ ID NO:110
<b>4</b> 5	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:110:
	Val Thr Ala Pro Lys Ser Ser Lys Gly Gly Ser Ser Ala Glu Lys Lys 1 5 10 15
55	

	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:118:
	Gln Thr Ser Val Arg Leu Gly Gln Gly Ser Ser Asp Pro Glu Lys Lys 1 5 10 15
10	(119) INFORMATION FOR SEQ ID NO:119
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:119:
20	Lys Thr Ser Leu Arg Pro Trp Lys Gly Ser Ser Asp Ser Asp Lys Lys  1 10 15
25	(120) INFORMATION FOR SEQ ID NO:120
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
00	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:120:
35	Gln Thr Asp Val Thr Gln Gly Gln Gly Ser Ser Gln Pro Glu Lys Lys 1 5 10 15
	(121) INFORMATION FOR SEQ ID NO:121
40	<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 16 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul>
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:121:
50	Gln Thr Ala Val Ser Gln Gly Gln Gly Ser Ser Gln Ser Glu Lys Lys 1 5 10 15
	(122) INFORMATION FOR SEQ ID NO:122
55	(i) SEQUENCE CHARACTERISTICS:

5	(A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:122:
10	Leu Thr Ala Pro Arg Thr Asn Arg Gly Ser Ser Asp Ser Glu Lys Lys 1 5 10 15
15	(123) INFORMATION FOR SEQ ID NO:123
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:123:
25	Val Thr Ala Pro Ser Ser His Arg Gly Ser Ser Asp Thr Glu Lys Lys 1 10 15
	(124) INFORMATION FOR SEQ ID NO:124
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:124:
40	Leu Leu Ser Leu Ser Pro Leu Lys Gly Asp Ser Asp Pro Glu Lys Val 1 5 10 15
	(125) INFORMATION FOR SEQ ID NO:125
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:125:
	Val Thr Ala Pr Thr Pro Asp Thr Gly Ala Ile Lys Thr Glu Lys Leu 1 5 10 15
56	

	(126) INFORMATION FOR SEQ ID NO:126
5	<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 16 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul>
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:126:
15	Val Thr Ile Pro Thr Pro Asp Thr Gly Ala Ile Lys Thr Glu Lys Leu  1 5 10 15
	(127) INFORMATION FOR SEQ ID NO:127
20	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:127:
	Ala Val Ser Pro Thr Pro Asp Thr Gly Ala Ile Lys Thr Glu Lys Leu  1 5 10 15
30	(128) INFORMATION FOR SEQ ID NO:128
35	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:128:
40	Ala Val Ser Pro Thr Pro Asp Thr Gly Ala Ile Lys Thr Glu Lys Leu 1 5 10 15
45	(129) INFORMATION FOR SEQ ID NO:129
	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:129:

5	1 5 10 11 15 Asp thr Gly Val He Lys Thr Glu Lys Leu 1 5 10 15
	(130) INFORMATION FOR SEQ ID NO:130
10	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:130:
20	Ala Val Ser Pro Thr Pro Asp Thr Gly Ala Ile Lys Thr Glu Pro Ser 1 5 10 15
	(131) INFORMATION FOR SEQ ID NO:131
25	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:131:
	Tyr Leu Pro Pro Thr Pro Gly Val Ile Arg Ser Thr Ala Met Lys Leu 1 5 10 15
35	(132) INFORMATION FOR SEQ ID NO:132
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid
40	(C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
45	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:132:
	Tyr Leu Pro Pro Thr Pro Gly Val Ile Arg Ser Thr Ala Met Arg Leu 1 5 10 15
50	(133) INFORMATION FOR SEQ ID NO:133
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
55	

	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:133:
	Tyr Leu Pro Pro Thr Pro Gly Leu Ile Arg Ser Thr Ser Met Lys Leu 1 5 10 15
10	(134) INFORMATION FOR SEQ ID NO:134
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:134:
20	Tyr Leu Pro Pro Thr Pro Gly Leu Ile Arg Ser Thr Ser Val Lys Leu 1 5 10 15
25	(135) INFORMATION FOR SEQ ID NO:135
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:135:
35	Tyr Leu Pro Pro Thr Pro Gly Val Ile Arg Ser Thr Ala Glu Lys Leu 1 5 10 15
	(136) INFORMATION FOR SEQ ID NO:136
40	<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 16 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul>
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:136:
50	Tyr Leu Pro Pro Thr Pro Gly Val Ile Arg Ser Thr Ala Gly Lys Leu 1 5 10 15
	(137) INFORMATION FOR SEQ ID NO:137
55	(i) SEQUENCE CHARACTERISTICS:

5	(A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: peptide	
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:137:	
10	Tyr Leu Pro Ala Thr Pro Gly Val Val Arg Ser Ser Ala Gly Met Leu 1 5 10 15	u
15	(138) INFORMATION FOR SEQ ID NO:138	
20	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: peptide	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:138:	
25	Ser Leu Pro Pro Ser Pro Gly Lys Val Arg Ser Thr Ala Glu Lys Let 1 5 10 15	u
	(139) INFORMATION FOR SEQ ID NO:139	
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear	
35	(ii) MOLECULE TYPE: peptide	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:139:	
<b>4</b> 0	Ser Leu Pro Pro Ser Pro Gly Lys Val Arg Ser Thr Ala Asn Lys Let 1 5 10 15	u.
	(140) INFORMATION FOR SEQ ID NO:140	
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: peptide	
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:140:	
	Ser Leu Pro Pro Arg Pro Gly Lys Val Arg Ser Ser Ser Glu Lys Le 1 5 10 15	u
55		

	(141) INFORMATION FOR SEQ ID NO:141
5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:141:
15	Ser Leu Pro Pro Arg Pro Gly Lys Val Arg Ser Ser Ser Asp Lys Leu  1 10 15
	(142) INFORMATION FOR SEQ ID NO:142
20	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
25	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:142:
30	Ser Leu Pro Pro Arg Pro Gly Arg Val Arg Ser Ser Ser Glu Lys Leu 1 5 10 15
	(143) INFORMATION FOR SEQ ID NO:143
35	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:143:
	Ser Leu_Pro Pro Arg Pro Gly Lys Val Arg Ser Ser Ser Glu Gln Leu 1 5 10 15
45	(144) INFORMATION FOR SEQ ID NO:144
50	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
E E	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:144:

_	1 5 10 15 FIG ALG PLO GLY LYS VAL ANG SET SET SET GLU THE Leu
5	(145) INFORMATION FOR SEQ ID NO:145
10	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:145:
	Ser Leu Pro Pro Lys Pro Gly Lys Ile Arg Ser Ser Thr Gly Lys Leu 1 5 10 15
20	(146) INFORMATION FOR SEQ ID NO:146
25	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:146: Ser Leu Pro Pro Lys Pro Gly Arg Ile Arg Ser Ser Thr Gly Lys Leu
	1 5 10 15
35	(147) INFORMATION FOR SEQ ID NO:147
	<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 16 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul>
40	( <u>i</u> i) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:147:
45	Ser Leu Pro Pro Lys Pro Gly Lys Ile Arg Ser Ser Thr Gly Gln Leu 1 5 10 15
	(148) INFORMATION FOR SEQ ID NO:148
50	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
55	

	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:148:
	Ser Leu Pro Pro Glu Pro Gly Lys Ile Arg Ser Ser Thr Gly Arg Leu 1 5 10 15
10	(149) INFORMATION FOR SEQ ID NO:149
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:149:
20	Ser Leu Ala Pro Ser Pro Gly Lys Ile Arg Ser Thr Ala Glu Lys Leu 1 5 10 15
25	(150) INFORMATION FOR SEQ ID NO:150
	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
30	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:150:
35	Ser Leu Pro Pro Arg Pro Gly Lys Ile Arg Ser Ser Thr Gly Asn Val 1 5 10 15
	(151) INFORMATION FOR SEQ ID NO:151
40	(i) SEQUENCE CHARACTERISTICS: - (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:151:
50	Ser Leu Arg Pro Ser Pro Gly Lys Val Arg Ser Thr Ala Glu Lys Leu 1 5 10 15
	(152) INFORMATION FOR SEQ ID NO:152
55	(i) SEQUENCE CHARACTERISTICS:

5	<ul><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:152:
	Ser Leu Arg Pro Ser Pro Gly Lys Val Arg Ser Thr Ala Asp Lys Leu 1 5 10 15
15	(153) INFORMATION FOR SEQ ID NO:153
20	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:153:
25	Ser Leu Arg Pro Ser Pro Gly Lys Val Arg Ser Thr Ala Glu Asn Leu 1 5 10 15
	(154) INFORMATION FOR SEQ ID NO:154
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:154:
<b>4</b> 0	Ser Leu Arg Pro Ser Pro Gly Lys Val Arg Ser Ala Val Glu Lys Leu 1 5 10 15
	(155) INFORMATION FOR SEQ ID NO:155
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 15 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:155:
	Ser Leu Pro Pro Arg Pro Gly Lys Arg Ser Ser Ala Glu Lys Leu 1 5 10 15

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	(156) INFORMATION FOR SEQ ID NO:156
5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:156:
15	Ser Leu Ala Pro Ser Pro Gly Lys Val Arg Ser Thr Val Glu Arg Leu 1 5 10 15
	(157) INFORMATION FOR SEQ ID NO:157
20	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:157:
	Ser Leu Ala Pro Ser Pro Asp Lys Ile Arg Ser Thr Pro Asp Lys Leu 1 5 10 15
30	(158) INFORMATION FOR SEQ ID NO:158
35	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid
	(C) TOPOLOGY: linear
40	(C) TOPOLOGY: linear
40	(C) TOPOLOGY: linear  (ii) MOLECULE TYPE: peptide
<b>40</b> <b>45</b>	(C) TOPOLOGY: linear  (ii) MOLECULE TYPE: peptide  (xi) SEQUENCE DESCRIPTION: SEQ ID NO:158:  Ser Leu_Ala Leu Ser Pro Gly Lys Val Arg Ser Thr Ala Glu Lys Leu
<b>4</b> 5	(C) TOPOLOGY: linear  (ii) MOLECULE TYPE: peptide  (xi) SEQUENCE DESCRIPTION: SEQ ID NO:158:  Ser Leu_Ala Leu Ser Pro Gly Lys Val Arg Ser Thr Ala Glu Lys Leu  1 5 10 15
	(ii) MOLECULE TYPE: peptide  (xi) SEQUENCE DESCRIPTION: SEQ ID NO:158:  Ser Leu_Ala Leu Ser Pro Gly Lys Val Arg Ser Thr Ala Glu Lys Leu 1 5 10 15  (159) INFORMATION FOR SEQ ID NO:159  (i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids (B) TYPE: amino acid
<b>4</b> 5	(ii) MOLECULE TYPE: peptide  (xi) SEQUENCE DESCRIPTION: SEQ ID NO:158:  Ser Leu_Ala Leu Ser Pro Gly Lys Val Arg Ser Thr Ala Glu Lys Leu  1 5 10 15  (159) INFORMATION FOR SEQ ID NO:159  (i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear

5	Ser Leu Pro Leu Ser Ala Gly Lys Val Arg Ser Thr Ala Glu Lys Leu 1 5 10 15
	(160) INFORMATION FOR SEQ ID NO:160
10	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:160:
	Ser Leu Ala Pro Ser Pro Gly Lys Val Arg Ser Thr Ala Glu Tyr Leu 1 5 10 15
20	(161) INFORMATION FOR SEQ ID NO:161
25	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:161:
30	Ser Leu Pro Leu Thr Pro Gly Leu Ile Arg Ser Thr Ala Glu Lys Leu 1 5 10 15
35	(162) INFORMATION FOR SEQ ID NO:162
40	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
40	(ii) MOLECULE TYPE: peptide
	(#i) SEQUENCE DESCRIPTION: SEQ ID NO:162:
<b>4</b> 5	Ser Leu Pro Leu Thr Pro Arg Val Ile Arg Ser Thr Ala Glu Lys Leu 1 5 10 15
	(163) INFORMATION FOR SEQ ID NO:163
50	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
55	

	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:163:
	Phe Leu His Pro Thr Pro Gly Thr Asp Ser Ser Ser Thr Glu Lys Leu 1 5 10 15
10	(164) INFORMATION FOR SEQ ID NO:164
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:164:
20	Phe Leu Leu Pro Thr Pro Gly Thr Asp Ser Ser Ser Thr Glu Arg Leu 1 5 10 15
25	(165) INFORMATION FOR SEQ ID NO:165
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:165:
35	Phe Leu His Pro Thr Arg Val Thr Asp Ser Ser Ser Thr Glu Lys Leu 1 5 10 15
•	(166) INFORMATION FOR SEQ ID NO:166
40	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:166:
50	Leu Leu Pro Pro Thr Pro Gly Thr Asn Ser Ser Ser Asn Asp Lys Leu 1 5 10 15
	(167) INFORMATION FOR SEQ ID NO:167
55	(i) SEQUENCE CHARACTERISTICS:

5	(B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:167:
10	Val Leu Pro Leu Ser Pro His Arg Ile Arg Ser Glu Ser Glu Asn Leu 1 5 10 15
15	(168) INFORMATION FOR SEQ ID NO:168
	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:168:
25	Ser Leu Ala Pro Ser Pro Ala Lys Phe Arg Ser Thr Ala Glu Arg Asp 1 5 10 15
	(169) INFORMATION FOR SEQ ID NO:169
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:169:
40	Val Thr Ala Pro Arg Pro Gly Arg Ile Arg Ser Asp Pro Glu Lys Lys 1 5 10 15
	(170) INFORMATION FOR SEQ ID NO:170
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:170:
	Val Thr Ala Pro Arg Pro Gly Arg Val Arg Ser Asp Pro Glu Lys Lys 1 5 10 15
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	(171) INFORMATION FOR SEQ ID NO:171
5	<ul> <li>(i) SEQUENCE CHARACTERISTICS:         <ul> <li>(A) LENGTH: 16 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul> </li> </ul>
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:171:
15	Val Thr Gly Pro Arg Pro Gly Arg Ile Arg Ser Asp Pro Glu Lys Lys 1 5 10 15
	(172) INFORMATION FOR SEQ ID NO:172
20	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:172:
	Val Thr Gly Pro Arg Pro Gly Arg Ile Arg Ser Asp Pro Asp Lys Lys 1 10 15
30	(173) INFORMATION FOR SEQ ID NO:173
35	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:173:
	Val Thr Gly Pro Arg Pro Gly Arg Val Arg Ser Asp Pro Glu Lys Lys 1 5 10 15
45	(174) INFORMATION FOR SEQ ID NO:174
50	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
~	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:174:
55	(112) 22 <b>3</b> 021102 222122222222

5	Val Thr Gly Pro Arg Pro Gly Arg Ile Arg Ser Asp Pro Xaa Lys Lys  1 10 15
	(175) INFORMATION FOR SEQ ID NO:175
10	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:175:
	Val Thr Ala Pro Arg Pro Gly Arg Ile Arg Ser Glu Ser Glu Arg Lys 1 5 10 15
20	(176) INFORMATION FOR SEQ ID NO:176
. 25	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:176:
30	Val Thr Gly Pro Ser Arg Gly Arg Ile Arg Ser Asp Pro Glu Lys Lys 1 5 10 15
35	(177) INFORMATION FOR SEQ ID NO:177
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
40	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:177:
<b>4</b> 5	Val Thr Val Pro Arg Pro Ser Arg Ile Arg Ser Glu Ser Glu Arg Lys 1 5 10 15
	(178) INFORMATION FOR SEQ ID NO:178
50	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear

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	(II) MODECODE III . populac
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:178:
	Val Thr Ala Pro Gly Pro Gly Arg Ile Arg Ser Glu Ser Glu Arg Lys  1 10 15
10	(179) INFORMATION FOR SEQ ID NO:179
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:179:
20	Gln Thr Ser Val Arg Pro Gly Arg Val Arg Ser Asp Pro Glu Arg Lys  1 10 15
25	(180) INFORMATION FOR SEQ ID NO:180
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
30	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:180:
35	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Asp Pro Glu Arg Lys 1 5 10 15
	(181) INFORMATION FOR SEQ ID NO:181
40	<ul> <li>(i) SEQUENCE CHARACTERISTICS:         <ul> <li>(A) LENGTH: 16 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul> </li> </ul>
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:181:
50	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Asp Pro Glu Lys Lys 1 5 10 15
	(182) INFORMATION FOR SEQ ID NO:182
55	(i) SEQUENCE CHARACTERISTICS:

5	(B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:182:
10	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Glu Pro Glu Lys Lys 1 5 10 15
	(183) INFORMATION FOR SEQ ID NO:183
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:183:
25	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Glu Pro Asp Lys Lys 1 5 10 15
	(184) INFORMATION FOR SEQ ID NO:184
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
35	(ii) MOLECULE TYPE: peptide
33	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:184:
	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ala Glu Pro Glu Lys Lys 1 10 15
40	(185) INFORMATION FOR SEQ ID NO:185
	(i) SEQUENCE CHARACTERISTICS:
45	(A) LENGTH: 16 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:185:
	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Asx Pro Glx Lys Lys 1 5 10 15

	(186) INFORMATION FOR SEQ ID NO:186
5	<ul> <li>(i) SEQUENCE CHARACTERISTICS:         <ul> <li>(A) LENGTH: 16 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul> </li> </ul>
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:186:
15	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Asp Pro Asx Lys Lys  1 5 10 15
	(187) INFORMATION FOR SEQ ID NO:187
20	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:187:
	Gln Thr Ser Val Arg Pro Gly Gln Val Arg Ser Asp Pro Glu Arg Lys 1 5 10 15
30	(188) INFORMATION FOR SEQ ID NO:188
35	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:188:
	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser His Pro Glu Lys Lys 1 5 10 15
45	(189) INFORMATION FOR SEQ ID NO:189
50	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:189:
55	

	1 5 10 19 ASH VAI AIG SET ASP PTO ASP Lys Lys
5	(190) INFORMATION FOR SEQ ID NO:190
10	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:190:
	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Asp Pro Glu Lys Thr 1 5 10 15
20	(191) INFORMATION FOR SEQ ID NO:191
25	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:191:
	Gln Thr Ser Val Arg Pro Gly Thr Val Arg Ser Glu Pro Glu Lys Lys 1 5 10 15
<b>35</b>	(192) INFORMATION FOR SEQ ID NO:192
40	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
<b>-</b>	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:192:
45	Gln Thr Ser Val Arg Pro Glu Lys Val Arg Ser Glu Pro Asp Lys Lys 1 5 10 15
	(193) INFORMATION FOR SEQ ID NO:193
50	<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 16 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul>
55	

	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:193:
	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Glu Ser Asp Lys Lys 1 5 10 15
10	(194) INFORMATION FOR SEQ ID NO:194
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:194:
20	Gln Thr Ser Val Arg Pro Gly Glu Val Arg Ser Glu Pro Asp Lys Lys 1 5 10 15
25	(195) INFORMATION FOR SEQ ID NO:195
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
00	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:195:
35	Gln Thr Ser Val Arg Pro Gly Asx Val Arg Ser Asx Pro Glx Arg Lys 1 5 10 15
	(196) INFORMATION FOR SEQ ID NO:196
40	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:196:
50	Gln Thr Ser Val Ser Pro Gly Lys Val Arg Ser Asp Pro Glu Lys Lys 1 5 10 15
	(197) INFORMATION FOR SEQ ID NO:197
55	(i) SEQUENCE CHARACTERISTICS:

5	(B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:197:
10	Gln Thr Ser Val Arg Pro Gly Lys Val Asn Ser Asp Pro Glu Lys Lys 1 5 10 15
15	(198) INFORMATION FOR SEQ ID NO:198
	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:198:
25	Gln Thr Ser Val Arg Pro Gly Lys Val Arg Ser Asp Pro Asp Thr Lys  1 10 15
	(199) INFORMATION FOR SEQ ID NO:199
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:199:
	Gln Thr Ser Val Arg Pro Lys Lys Val Arg Ser Asp Pro Glx Lys Lys  1 10 15
40	1 5 10 15
	(200) INFORMATION FOR SEQ ID NO:200
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:200:
	Gln Thr Ser Val Arg Pro Lys Lys Val Arg Phe Asp Pro Glu Lys Lys 1 5 10 15
55	

	(201) INFORMATION FOR SEQ ID NO:201
5	<ul> <li>(i) SEQUENCE CHARACTERISTICS:         <ul> <li>(A) LENGTH: 16 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul> </li> </ul>
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:201:
15	Gln Thr Ser Val Arg Ser Gly Lys Val Arg Ser Glu Pro Glu Thr Lys  1 5 10 15
	(202) INFORMATION FOR SEQ ID NO:202
20	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:202:
	Val Thr Asn Leu Arg Pro Gly Lys Val Arg Ser Asp Ala Glu Lys Lys 1 5 10 15
30	(203) INFORMATION FOR SEQ ID NO:203
35	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:203:
	Val Thr Asp Leu Arg Pro Gly Lys Val Arg Ser Asp Ala Glu Lys Lys 1 5 10 15
45	(204) INFORMATION FOR SEQ ID NO:204
50	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:204:
55	

5	1 5 10 Ser Val Ser Pro Gly Asn Ile Arg Ser Glu Ser Asp Lys Lys
	(205) INFORMATION FOR SEQ ID NO:205
10	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:205:
*	Lys Thr Ser Val Thr Pro Gly Lys Phe Arg Ser Glu Pro Glu Lys Lys 1 5 10 15
20	(206) INFORMATION FOR SEQ ID NO:206
25	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) WOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:206:
30	Val Thr Leu Leu Pro Pro Gly Arg Val Arg Ser Asp Ala Glu Lys Lys 1 5 10 15
35	(207) INFORMATION FOR SEQ ID NO:207
40	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
40	(11) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:207:
45	Val Thr Leu Leu Pro Pro Gly Glu Val Arg Ser Asp Ala Glu Lys Lys 1 5 10 15
	(208) INFORMATION FOR SEQ ID NO: 208
50	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
e e	

	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:208:
	Val Thr Leu Pro Pro Pro Gly Glx Val Arg Ser Asp Ala Glu Arg Lys 1 5 10 15
10	(209) INFORMATION FOR SEQ ID NO:209
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:209:
20	Val Thr Leu Pro Pro Pro Gly Glx Val Arg Ser Asx Ala Glx Asn Lys 1 5 10 15
25	(210) INFORMATION FOR SEQ ID NO:210
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:210:
35	Val Thr Leu Pro Pro Pro Gln Gln Val Arg Ser Asp Ala Glu Lys Lys 1 5 10 15
	(211) INFORMATION FOR SEQ ID NO:211
40	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:211:
50	Val Thr Leu Pro Pro Pro Gly Gln Val Thr Ser Asp Ala Glu Lys Lys 1 5 10 15
	(212) INFORMATION FOR SEQ ID NO:212
	(i) SEQUENCE CHARACTERISTICS:

5	<ul><li>(A) LENGTH: 16 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:212:
10	Val Thr Leu Pro Pro Ala Gly Gln Val Arg Ser Asp Ala Glu Lys Arg 1 5 10 15
15	(213) INFORMATION FOR SEQ ID NO:213
20	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 16 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:213:
25	Ala Leu Ser Pro Ser Ser Gly Gln Ser Ser Ser Ala Ser Glu Arg Leu 1 5 10 15
	(214) INFORMATION FOR SEQ ID NO:214
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:214:
40	Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ala 1 5 10 15
	Ser Arg Gly Asp Ser Gln Arg Pro Glu Ser 20 25
45	(215) INFORMATION FOR SEQ ID NO:215
50	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:215:
55	

5	1 5 10 15 15
	Ser Arg Gly Asp Ser Gln Arg Pro Glu Ser 20 25
10	(216) INFORMATION FOR SEQ ID NO:216
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:216:
20	Glu Lys Val Gly Gly Leu Gln Pro Gly Thr Gly Ala Pro Gly Lys Ala 1 5 10 15
25	Ser Arg Gly Asp Ser Gln Arg Pro Glu Ser 20 25
	(217) INFORMATION FOR SEQ ID NO:217
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:217:
	Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ala 1 5 10 15
40	Ser Lys Gly Asn Ser Gln Arg Ala Glu Ser
40	20
	(218) INFORMATION FOR SEQ ID NO:218
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
<i>3</i> 0	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:218:

108

5	1 5 10 17 The Gly Lys Ala
	Ser Lys Gly Asn Ser Gln Arg Ala Glu Ser 20 25
10	(219) INFORMATION FOR SEQ ID NO:219
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:219:
20	Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ala 1 5 10 15
25	Ser Lys Gly Thr Ser Gln Arg Ala Glu Ser 20 25
	(220) INFORMATION FOR SEQ ID NO:220
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:220:
	Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ala 1 5 10 15
40	Ser Ly® Gly Thr Ser Gln Arg Ala Glu Thr 20 25
	(221) INFORMATION FOR SEQ ID NO:221
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:221:

	Glu Lys Val Gly Gly Leu Lys Pro Gly Arg Gly Thr Pro Gly Lys Ala 1 5 10 15
5	
	Ser Lys Gly Thr Ser Gln Arg Ala Glu Thr 20 25
10	(222) INFORMATION FOR SEQ ID NO:222
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
,,,	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:222:
20	Glu Asn Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ala 1 5 10 15
25	Ser Lys Gly Thr Ser Gln Arg Ala Glu Thr 20 25
	(223) INFORMATION FOR SEQ ID NO:223
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:223:
	Glu Lys Val Gly Gly Leu Gln Ser Gly Arg Gly Thr Pro Gly Lys Ala 1 5 10 15
40	Ser Lys Gly Thr Ser Gln Arg Ala Glu Thr
	(224) INFORMATION FOR SEQ ID NO:224
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:224:

140

5	1 5 10 15 15 15 16 17 Leu Gin Ser Gly Arg Gly Thr Pro Gly Lys Ala
	Ser Lys Gly Thr Ser Gln Arg Ala Glu Ser 20 25
10	(225) INFORMATION FOR SEQ ID NO:225
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:225:
20	Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ala 1 5 10 15
25	Ser Lys Gly Ile Ser Gln Arg Ala Glu Arg 20 25
	(226) INFORMATION FOR SEQ ID NO:226
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:226:
	Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ser 1 5 10 15
40	Ala Lys Gly Asx Ser Glx Arg Ala Gln Ser
	(227) INFORMATION FOR SEQ ID NO:227
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:227:

5	Glu Lys Val Gly Gly Leu Gln Pro Gly Ser Gly Thr Pro Gly Lys Ala 1 5 10 15
	Ser Lys Gly Asn Ser Gln Arg Ala Glu Ser 20 25
10	(228) INFORMATION FOR SEQ ID NO:228
15	<ul> <li>(i) SEQUENCE CHARACTERISTICS:         <ul> <li>(A) LENGTH: 26 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul> </li> </ul>
	(ii) MOLECULE TYPE: peptide
00	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:228:
20	Glu Lys Val Gly Gly Leu Gln Pro Gly Ser Gly Thr Pro Gly Lys Ala 1 5 10 15
25	Ser Lys Gly Ser Ser Gln Arg Ala Glu Ser 20 25
	(229) INFORMATION FOR SEQ ID NO:229
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:229:
40	Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Arg Lys Ala 1 5 10 15
40	Ser Lys Gly Asn Ser Gln Arg Ala Glu Ser 20 25
<b>4</b> 5	(230) INFORMATION FOR SEQ ID NO:230
-,0	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:230:

5	1 5 10 Lys Met Gly Ash Leu Gln Pro Gly Ser Gly Thr Pro Gly Lys Ala
	Ser Lys Gly Asn Ser Gln Arg Pro Asp Ser 20 25
10	(231) INFORMATION FOR SEQ ID NO:231
	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:231:
20	Glu Lys Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
25	Ser Lys Gly Asn Ala Arg Arg Ser Glu Thr 20 25
	(232) INFORMATION FOR SEQ ID NO:232
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:232:
-	Glu Lys Val Gly Gly Leu Lys Pro Gly Lys Gly Ala Pro Glu Lys Asp 1 10 15
40	Ser Lys Gly Asn Ala Arg Arg Ser Glu Thr 20 25
	(233) INFORMATION FOR SEQ ID NO:233
<b>4</b> 5	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:233:

5	1 5 10 Lys Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Arg Asp
	Ser Lys Gly Asn Ala Arg Arg Ser Glu Thr 20 25
10	(234) INFORMATION FOR SEQ ID NO:234
15	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:234:
20	Asp Lys Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
25	Ser Lys Gly Asn Ala Lys Arg Ser Glu Thr 20 25
	(235) INFORMATION FOR SEQ ID NO:235
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:235:
	Asp Lys Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
40	Ser Lys Gly Asn Ala Lys Lys Ser Glu Thr
	(236) INFORMATION FOR SEQ ID NO:236
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:236:

5	Asp Lys Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Asp Lys Asp 1 5 10 15
	Asn Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
10	(237) INFORMATION FOR SEQ ID NO:237
	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:237:
20	Glu Lys Val Gly Gly Leu Thr Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
25	Ser Lys Gly Asn Gly Arg Arg Ser Glu Thr 20 25
	(238) INFORMATION FOR SEQ ID NO:238
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:238:
•	Glu Met Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
40	Ser Lys Gly Asn Asp Arg Arg Ser Glu Thr
	(239) INFORMATION FOR SEQ ID NO:239
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:239:

	Glu Met Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 1 15
5	Ser Lys Gly Asn Asp Lys Arg Ser Glu Thr
	(240) INFORMATION FOR SEQ ID NO:240
10	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:240:
20	Glu Met Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 10 15
	Ser Lys Gly Asn Ala Lys Arg Ser Glu Thr 20 25
25	(241) INFORMATION FOR SEQ ID NO:241
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:241:
35	Glu Gln Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
40	Ser Lys Gly Asn Ala Lys Lys Ser Glu Thr
	(242) INFORMATION FOR SEQ ID NO:242
<b>4</b> 5	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
<b>5</b> 0	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:242:

5	Glu Gln Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 10 15
	Thr Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
10	(243) INFORMATION FOR SEQ ID NO:243
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:243:
20	Glu Gln Val Gly Gly Leu Lys Pro Gly Lys Gly Ala Pro Glu Lys Asp 1 5 10 15
25	Thr Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
	(244) INFORMATION FOR SEQ ID NO: 244
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 244:
	Glu Lys Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 10 15
40	Ser Lyg Gly Asn Ala Lys Lys Ser Glu Thr 20 25
	(245) INFORMATION FOR SEQ ID NO:245
<b>4</b> 5	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:245:

5	Glu Lys Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
J	Thr Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
10	(246) INFORMATION FOR SEQ ID NO:246
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:246:
20	Glu Lys Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Glu Lys Asp 1 5 10 15
25	Thr Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
	(247) INFORMATION FOR SEQ ID NO:247
3 <i>0</i>	<ul> <li>(i) SEQUENCE CHARACTERISTICS:         <ul> <li>(A) LENGTH: 26 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul> </li> </ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 247:
<i>:</i>	Glu Lys Val Gly Gly Leu Gln Pro Gly Lys Gly Ser Pro Glu Lys Asp 1 10 15
40	Ser Lys Gly Asn Ala Lys Lys Ser Glu Thr
	(248) INFORMATION FOR SEQ ID NO:248
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:248:

5	1 5 10 10 17 15 15 15 15 15 15 15 15 15 15 15 15 15
	Ser Lys Gly Asn Ala Lys Gln Ser Glu Thr 20 25
10	(249) INFORMATION FOR SEQ ID NO:249
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:249:
20	Glu Gln Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Pro Asp Lys Asp 1 5 10 15
25	Ser Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
	(250) INFORMATION FOR SEQ ID NO:250
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:250:
	Glu Lys Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
40	Ser Lys Gly Asn Ala Glu Lys Ser Glu Thr 20 25
	(251) INFORMATION FOR SEQ ID NO:251
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
<i>3</i> 0	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:251:

5	Glu Gln Val Gly Asp Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 10 15
3	Thr Lys Gly Asn Ala Arg Arg Ser Glu Thr 20 25
10	(252) INFORMATION FOR SEQ ID NO:252
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:252:
20	Glu Asn Val Gly Asp Leu Lys Pro Gly Lys Gly Ala Pro Glu Lys Asp 1 5 10 15
25	Ser Lys Gly Asn Ala Arg Arg Ser Glu Thr 20 25
	(253) INFORMATION FOR SEQ ID NO:253
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:253:
· .	Glu Gln Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Ser Asp Lys Asp 10 15
40	Ser Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
	(254) INFORMATION FOR SEQ ID NO:254
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:254:

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5	1 5 10 17 Pro Glu Lys Asp
	Ser Lys Gly Asn Ala Lys Lys Ser Gly Thr 20 25
10	(255) INFORMATION FOR SEQ ID NO:255
15	<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 26 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:255:
20	Asp Gln Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
25	Thr Lys Gly Asn Pro Lys Arg Ser Glu Thr 20 25
	(256) INFORMATION FOR SEQ ID NO:256
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:256:
	Asp Gln Val Gly Gly Leu Gln Pro Gly Gln Gly Thr Pro Glu Lys Asn 1 5 10 15
40	Thr Lys Gly Asn Pro Lys Arg Ser Asp Thr
	(257) INFORMATION FOR SEQ ID NO:257
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:257:

5	Giu Lys Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Ser Glu Lys Asp 1 5 10 15
•	Ile Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
10	(258) INFORMATION FOR SEQ ID NO:258
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:258:
20	Asp Lys Val Gly Gly Leu Lys Pro Gly Lys Arg Thr Pro Glu Lys Asp 1 5 10 15
25	Asn Lys Gly Asn Ala Lys Lys Ser Glu Thr 20 25
20	(259) INFORMATION FOR SEQ ID NO:259
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:259:
	Asp Lys Val Gly Gly Leu Lys Leu Gly Lys Gly Thr Pro Glu Lys Asp 1 10 15
40	Thr Lys Gly Asn Ala Lys Lys Ser Glu Thr
	(260) INFORMATION FOR SEQ ID NO:260
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:260:

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5	1 5 10 15
	Ser Lys Gly Asn Ala Asn Thr Ser Glu Thr 20 25
10	(261) INFORMATION FOR SEQ ID NO:261
15	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:261:
20	Glu His Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
25	Ser Lys Gly Asn Ala Gly Arg Ser Glu Thr 20 25
	(262) INFORMATION FOR SEQ ID NO:262
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:262:
	Glu Gln Val Gly Gly Leu Gln Pro Gly Asn Gly Thr Pro Glu Lys Asp 1 5 10 15
40	Thr The Gly Asn Ala Lys Arg Ser Glu Thr
	(263) INFORMATION FOR SEQ ID NO: 263
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 263:

5	1 5 10 15 Glu Lys Glu
	Ser Lys Gly Asp Ser Lys Arg Ala Glu Thr 20 25
10	(264) INFORMATION FOR SEQ ID NO:264
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:264:
20	Glu Lys Glu Gly Gly Leu Gln Pro Gly Lys Gly Thr Pro Glu Lys Glu 1 5 10 15
25	Ser Lys Gly Asp Ser Lys Arg Pro Glu Thr 20 25
	(265) INFORMATION FOR SEQ ID NO:265
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:265:
	Glu Lys Glu Gly Gly Leu Gln Pro Gly Lys Gly Ser Pro Glu Lys Glu 1 5 10 15
40	Ser Lys Gly Asp Ser Lys Arg Ala Glu Thr
	(266) INFORMATION FOR SEQ ID NO:266
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
<i></i>	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:266:

5	1 5 10 15 Asp Gly Gly Led Gln Pro Gly Lys Gly Thr Pro Glu Lys Asp
	Ser Lys Gly Asp Ser Lys Arg Val Glu Met 20 25
10	(267) INFORMATION FOR SEQ ID NO:267
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:267:
20	Glu Gln Val Gly Gly Leu Lys Pro Gly Arg Gly Thr Pro Glu Lys Asp 1 5 10 15
25	Thr Thr Gly Asp Ala Gln Arg Ser Glu Thr 20 25
	(268) INFORMATION FOR SEQ ID NO:268
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:268:
	Glu Gln Val Gly Gly Leu Lys Pro Gly Arg Gly Thr Pro Glu Lys Asp 1 5 10 15
40	Thr Thr Gly Asn Ala Lys Gly Ser Glu Thr 20 25
	(269) INFORMATION FOR SEQ ID NO: 269
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
<b>5</b> 0	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:269:

5	Glu Lys Val Gly Gly Ser Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
	Ser Lys Gly Asn Ala Lys Thr Ser Glu Thr 20 25
10	(270) INFORMATION FOR SEQ ID NO:270
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:270:
20	Ser Asp Gln Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
25	Thr Lys Gly Asn Ala Arg Arg Ser Glu Ser 20 25
	(271) INFORMATION FOR SEQ ID NO:271
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:271:
	Glu Lys Ile Gly Gly Leu Gln Pro Gly Lys Gly Asp Pro Gly Lys Pro 1 5 10 15
40	Ser Lys Asp Asn Ala Lys Arg Ser Glu Thr
	(272) INFORMATION FOR SEQ ID NO:272
<b>4</b> 5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:272:

5	Glu Lys Leu Gly Gly Leu Gln Pro Gly Lys Gly Asp Pro Gly Lys Pro 1 5 10 15
	Ser Lys Asp Asn Ala Lys Arg Ser Glu Thr 20 25
10	(273) INFORMATION FOR SEQ ID NO:273
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:273:
20	Glu Lys Leu Gly Gly Leu Gln Pro Gly Lys Gly Asp Pro Gly Lys Pro 1 5 10 15
25	Phe Lys Asp Asn Ala Lys Arg Ser Glu Thr 20 25
	(274) INFORMATION FOR SEQ ID NO:274
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:274:
	Glu Lys Leu Gly Gly Leu Gln Pro Gly Lys Gly Asp Pro Gly Lys Leu 1 5 10 15
40	Met Lys Glu Asn Ala Lys Arg Ser Glu Thr 20 25
	(275) INFORMATION FOR SEQ ID NO: 275
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:275:

5	1 5 10 15 15
	Lys Xaa Glu Asn Ala Lys Arg Pro Glu Thr 20 25
10	(276) INFORMATION FOR SEQ ID NO:276
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:276:
20	Glu Lys Leu Gly Gly Leu Gln Pro Gly Asn Gly Asp Leu Gly Lys Pro 1 5 10 15
25	Ser Lys Asp Asn Ala Lys Arg Ser Glu Thr 20 25
	(277) INFORMATION FOR SEQ ID NO:277
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:277:
	Glu Lys Leu Gly Pro Leu Gln Leu Gly Lys Gly Asp Pro Gly Lys Pro 1 5 10 15
40	Ser Lys Asp Asp Ala Lys Arg Ser Glu Thr
	(278) INFORMATION FOR SEQ ID NO:278
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:278:

5	1 5 10 15
	Ser Lys Asp Asn Asp Lys Arg Ser Glu Thr 20 25
10	(279) INFORMATION FOR SEQ ID NO:279
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15	(11) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:279:
20	Glu Gln Leu Gly Gly Leu Gln Pro Gly Gly Gly Thr Pro Gly Lys Ala 1 5 10 15
25	Ser Lys Asp Asn Asp Lys Arg Ser Glu Thr 20 25
	(280) INFORMATION FOR SEQ ID NO:280
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:280:
	Glu Gln Val Gly Gly Leu Lys Ala Arg Lys Gly Thr Pro Glu Lys Asp 1 5 10 15
40	Thr Thr Gly Asn Ala Lys Arg Ser Glu Thr 20 25
	(281) INFORMATION FOR SEQ ID NO:281
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:281:

5	Glu Met Val Gly Val Leu Glu Pro Gly Lys Gly Thr Pro Glu Lys Arg 1 5 10 15
10	Gln Glu Gly Asn Ala Lys Arg Ser Glu Thr 20 25
	(282) INFORMATION FOR SEQ ID NO:282
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:282:
	Glu Gln Val Gly Gly Leu Gln Pro Lys Lys Gly Ser Pro Gly Lys Asp 1 10 15
25	Ser Lys Asp Asp Ser Gln Lys Thr Glu Thr
	(283) INFORMATION FOR SEQ ID NO:283
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:283:
	Glu Gln Val Gly Gly Leu Gln Pro Lys Lys Gly Ser Pro Gly Lys Asp 1 5 10 15
40	Ser Lys Asp Asp Ser Gln Lys Thr Glu Arg
	(284) INFORMATION FOR SEQ ID NO:284
<b>4</b> 5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:284:

5	1 5 10 11 15 11 15 11 15 15 15 10 10 15
	Asp Lys Gly Thr Ser Ala Arg Asn Asp Thr 20 25
10	(285) INFORMATION FOR SEQ ID NO:285
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
73	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:285:
20	Gln Gln Val Pro Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Lys Asp 1 5 10 15
25	Asp Lys Gly Thr Ser Ala Lys Asn Glu Thr 20 25
25	(286) INFORMATION FOR SEQ ID NO:286
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:286:
	Gln Gln Val Pro Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Lys Asp 1 10 15
40	Asp Lys Gly Thr Ser Ala Lys Asn Glu Met
	(287) INFORMATION FOR SEQ ID NO: 287
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:287:

5	1 5 10 Ser Pro Gly Leu Lys Pro Gly Lys Gly Ser Pro Gly Gln Glu 1 5 10 15
	Lys Lys Gly Thr Ser Ser Thr Ser Glu Thr 20 25
10	(288) INFORMATION FOR SEQ ID NO:288
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
,	(ii) MOLECULE TYPE: peptide
•	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:288:
20	Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
25	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
	(289) INFORMATION FOR SEQ ID NO:289
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:289:
	Glu Gln Gln Pro Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser
	(290) INFORMATION FOR SEQ ID NO:290
<b>4</b> 5	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:290:

5	Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
	Lys Lys Gly Lys Ser Ser Ala Ser Glu Ser 20 25
10	(291) INFORMATION FOR SEQ ID NO:291
	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:291:
20	Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Lys Gln 1 5 10 15
25	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
	(292) INFORMATION FOR SEQ ID NO:292
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:292:
	Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Thr His Gly Lys Gln 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
	(293) INFORMATION FOR SEQ ID NO:293
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:293:

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5	Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Ser His Gly Lys Gln 1 5 10 15
	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
10	(294) INFORMATION FOR SEQ ID NO:294
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:294:
20	Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Ser His Gly Lys Gln 1 5 10 15
25	Lys Lys Gly Lys Ser Ser Ala Ser Glu Ser 20 25
	(295) INFORMATION FOR SEQ ID NO:295
30	<ul> <li>(i) SEQUENCE CHARACTERISTICS:         <ul> <li>(A) LENGTH: 26 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul> </li> </ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:295: Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Thr His Gly Lys Gln
	1 5 10 15
40	Lys Lys Gly Lys Ser Ser Thr Phe Glu Ser
	(296) INFORMATION FOR SEQ ID NO:296
<b>4</b> 5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:296:

5	Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Thr His Gly Ly 1 5 10	ys Gln 15
10	Lys Gln Gly Lys Ser Ser Thr Phe Glu Ser 20 25	
	(297) INFORMATION FOR SEQ ID NO:297	
15	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: peptide	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:297:	
	Glu Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Thr His Gly Ly 1	ys Glu 15
25	Lys Lys Asp Lys Ser Ser Thr Ser Glu Ser 20 25	
	(298) INFORMATION FOR SEQ ID NO:298	
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: peptide	
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:298:	
	Glu Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Ser His Gly Ly 1 5 10	ys Gln 15
40	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser	
	(299) INFORMATION FOR SEQ ID NO:299	
<b>4</b> 5	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear	
50	(ii) MOLECULE TYPE: peptide	
JU	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:299:	

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5	Glu Gin Gin Pro Glu Leu Lys Pro Gly Lys Gly Thr His Gly Lys Gln 1 10 15
	Lys Lys Ser Asn Ser Ser Thr Ser Glu Ser 20 25
10	(300) INFORMATION FOR SEQ ID NO:300
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:300:
20	Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Ala Pro Gly Gln Glu 1 5 10 15
25	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
	(301) INFORMATION FOR SEQ ID NO:301
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:301:
	Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Ala Pro Gly Gln Glu 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Thr Ser Asp Ser
	(302) INFORMATION FOR SEQ ID NO:302
<b>4</b> 5	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:302:

5	Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Val Pro Gly Gln Glu 1 5 10 15
10	Lys Lys Gly Lys Ser Ser Thr Ser Asp Ser 20 25  (303) INFORMATION FOR SEQ ID NO:303
15	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	<pre>(ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:303:</pre>
20	Gln Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Ala Pro Gly Lys Gly 1 5 10 15
25	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
	(304) INFORMATION FOR SEQ ID NO:304
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:304:
	Gln Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Ala Pro Gly Lys Gly 1 5 10 15
40	Lys Lys Asp Lys Ser Ser Thr Ser Glu Ser 20 25
	(305) INFORMATION FOR SEQ ID NO:305
<b>4</b> 5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:305:

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5	Glu Gln Gln Pro Glu Ala Lys Pro Gly Lys Gly Thr His Gly Lys Gln 1 10 15
	Lys Lys Gly Lys Ser Ser Thr Ser Asp Ser 20 25
10	(306) INFORMATION FOR SEQ ID NO:306
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:306:
20	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr His Gly Lys Glu 1 5 10 15
25	Lys Lys Asp Lys Ser Ser Thr Ser Asp Ser 20 25
	(307) INFORMATION FOR SEQ ID NO:307
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:307:
	Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Ala Pro Gly Gln Gly 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser
	(308) INFORMATION FOR SEQ ID NO:308
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:308:

5	of Gin Gin Ala Glu Leu Lys Pro Gly Arg Gly Thr Pro Gly Gln Glu  1 10 15
10	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
	(309) INFORMATION FOR SEQ ID NO:309
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:309:
20	Glu Gln Gln Ala Glu Leu Arg Ala Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
25	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
	(310) INFORMATION FOR SEQ ID NO:310
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:310:
	Glu Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
<b>4</b> 0	Lys Lys Gly Thr Ser Ser Thr Ser Glu Ser
	(311) INFORMATION FOR SEQ ID NO: 311
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:311:

5	of the Gin Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly His Glu  1 10 15
	Lys Lys Gly Thr Ser Ser Thr Ser Glu Ser 20 25
10	(312) INFORMATION FOR SEQ ID NO:312
15	<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 26 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:312:
20	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly His Glu 1 5 10 15
25	Lys Lys Gly Thr Ser Ser Thr Ser Glu Ser 20 25
	(313) INFORMATION FOR SEQ ID NO:313
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:313:
	Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly His Glu 1 5 10 15
<b>4</b> 0	Asn Lys Gly Thr Ser Ser Thr Ser Glu Ser
	(314) INFORMATION FOR SEQ ID NO:314
<b>4</b> 5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
- <del>-</del>	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:314:

5	Gln Gln Gln Ala Glu Val Arg Pro Gly Lys Gly Thr Pro Gly His Glu 1 10 15
	Lys Lys Gly Thr Ser Ser Thr Ser Glu Ser 20 25
10	(315) INFORMATION FOR SEQ ID NO:315
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:315:
20	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly His Glu 1 5 10 15
25	Asn Lys Gly Thr Ser Ser Thr Ser Glu Ser 20 25
	(316) INFORMATION FOR SEQ ID NO:316
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:316:
	Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Ala Ser Glu Ser 20 25
	(317) INFORMATION FOR SEQ ID NO:317
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:317:

	His Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Gl 1 5 10 15	.n
5	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25	
10	(318) INFORMATION FOR SEQ ID NO:318	
	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>	•
15	(ii) MOLECULE TYPE: peptide	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:318:	
20	Glu Gln Gln Val Glu Leu Arg Ala Gly Lys Gly Thr Pro Gly Gln Gl 1 5 10 15	u
25	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25	
	(319) INFORMATION FOR SEQ ID NO:319	•
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>	
	(ii) MOLECULE TYPE: peptide	
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:319:	
	Glu Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15	u
40	Lys Gln Gly Thr Ser Ser Thr Ser Glu Ser	
	(320) INFORMATION FOR SEQ ID NO:320	
<b>1</b> 5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>	
50	(ii) MOLECULE TYPE: peptide	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:320:	

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5	1 5 15 15 15 15 15 15 15 15 15 15 15 15
	Asn Lys Gly Thr Ser Ser Thr Ser Glu Ser 20 25
10	(321) INFORMATION FOR SEQ ID NO:321
15	<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 26 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:321:
20	Gln Gln Gln Ala Glu Val Arg Pro Gly Lys Gly Thr Pro Gly His Glu 1 5 10 15
25	Lys Lys Gly Arg Ser Ser Thr Ser Glu Ser 20 25
	(322) INFORMATION FOR SEQ ID NO: 322
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:322:
	Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
40	Lys Lys Asp Lys Ser Ser Thr Ser Glu Ser
	(323) INFORMATION FOR SEQ ID NO:323
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:323:

5	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
	Lys Lys Asp Lys Ser Ser Thr Ser Glu Ser 20 25
10	(324) INFORMATION FOR SEQ ID NO: 324
15	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:324:
20	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
25	Lys Lys Asp Lys Ser Ser Thr Ser Asp Ser 20 25
	(325) INFORMATION FOR SEQ ID NO:325
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:325:
	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Ser Pro Gly Gln Gln 1 5 10 15
40	Lys Lys Asp Lys Ser Ser Thr Ser Glu Ser
	(326) INFORMATION FOR SEQ ID NO:326
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:326:

5	1 5 10 17 17 17 18 18 19 19 19 19 19 19 19 19 19 19 19 19 19
	Lys Lys Asn Lys Ser Ser Thr Ser Glu Ser 20 25
10	(327) INFORMATION FOR SEQ ID NO:327
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:327:
20	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
25	Asn Lys Asp Lys Ser Ser Thr Ser Glu Ser 20 25
	(328) INFORMATION FOR SEQ ID NO:328
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:328: Glu Gln Gln Ala Glu Leu Arg Ala Gly Lys Gly Ile Pro Gly Gln Glu
	1 5 10 15 15
40	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser
	(329) INFORMATION FOR SEQ ID NO:329
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
<b>5</b> 0	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:329:

5	1 5 10 15 15 15 16 16 16 16 16 16 16 16 16 16 16 16 16
	Lys Lys Ser Lys Ser Ser Thr Ser Glu Ser 20 25
10	(330) INFORMATION FOR SEQ ID NO:330
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:330:
20	Gln Gln Gln Ser Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
25	Lys Lys Ser Lys Ser Ser Thr Ser Glu Ser 20 25
	(331) INFORMATION FOR SEQ ID NO:331
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:331:
	Gln Gln Gln Thr Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
40	Lys Lys Ser Lys Ser Ser Thr Ser Glu Ser
	(332) INFORMATION FOR SEQ ID NO:332
<b>4</b> 5	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:332:

5	1 5 10 17 Pro Gly Gln Glu
	Arg Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
10	(333) INFORMATION FOR SEQ ID NO:333
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
,,	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:333:
20	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
25	Lys Lys Asp Lys Ser Ser Thr Phe Glu Ser 20 25
	(334) INFORMATION FOR SEQ ID NO:334
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
-	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:334:
	Glu Gln Gln Ala Glu Leu Arg Pro Gly Thr Gly Ala Pro Gly Gln Glu 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser
	(335) INFORMATION FOR SEQ ID NO:335
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
~ <b>~</b>	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:335:

5	Gln Gln Gln Pro Glu Val Arg Pro Gly Lys Gly Thr His Ala Lys Gln 1 5 10 15
	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
10	(336) INFORMATION FOR SEQ ID NO:336
15	<ul> <li>(i) SEQUENCE CHARACTERISTICS:         <ul> <li>(A) LENGTH: 26 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul> </li> </ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:336:
20	Gln Gln Gln Pro Glu Val Arg Pro Gly Lys Asp Thr His Ala Lys Gln 1 5 10 15
25	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
	(337) INFORMATION FOR SEQ ID NO:337
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:337:
	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Glu Gln Glu 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser
	(338) INFORMATION FOR SEQ ID NO:338
<b>4</b> 5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:338:

5	1 5 10 10 10 10 10 10 10 10 10 10 10 10 10
	Lys Lys Gly Arg Ser Ser Thr Ser Glu Ala 20 25
10	(339) INFORMATION FOR SEQ ID NO:339
15	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:339:
20	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Arg Glu 1 5 10 15
25	Lys Lys Ser Lys Pro Ser Thr Ser Glu Ser 20 25
	(340) INFORMATION FOR SEQ ID NO:340
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 340:
	Gln Gln Gln Ser Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Arg Glu 1 5 10 15
40	Lys Lys Ser Lys Pro Ser Thr Ser Glu Ser 20 25
	(341) INFORMATION FOR SEQ ID NO:341
<b>4</b> 5	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:341:

5	of the firm of the
	Lys Lys Asn Lys Pro Ser Thr Ser Glu Ser 20 25
10	(342) INFORMATION FOR SEQ ID NO:342
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:342:
20	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Arg Glu 1 5 10 15
25	Lys Lys Ser Thr Ser Ser Thr Ser Glu Ser 20 25
	(343) INFORMATION FOR SEQ ID NO:343
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:343:
	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
40	Lys Lys Ser Thr Ser Ser Thr Ser Asp Ser
	(344) INFORMATION FOR SEQ ID NO:344
<b>4</b> 5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 344:

Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Ile Gln Gln 10 5 Lys Lys Asp Lys Ser Ser Thr Ser Glu Ser (345) INFORMATION FOR SEQ ID NO:345 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:345: Gln Gln Gln Ala Glu Phe Lys Pro Gly Lys Gly Thr Pro Gly Arg Glu 20 His Arg Ser Lys Pro Ser Thr Ser Glu Ser 25 (346) INFORMATION FOR SEQ ID NO:346 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids
(B) TYPE: amino acid
(C) TOPOLOGY: linear 30 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:346: 35 Gln Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Ala Leu Gly Gln Glu Lys Lys Gly Lys Ser Ser Thr Ser Asp Ser 40 20 (347) INFORMATION FOR SEQ ID NO:347 (i) SEQUENCE CHARACTERISTICS: 45 (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:347:

Gln Gln Gln Pro Glu Val Lys Pro Gly Lys Gly Ala Pro Gly Lys Gly Asn Thr Asp Lys Ser Ser Thr Ser Glu Ser 20 (348) INFORMATION FOR SEQ ID NO:348 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear 15 (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:348: Glu Gln Gln Ala Glu Val Arg Ala Gly Lys Gly Ser Pro Gly Gln Glu 20 Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser 25 (349) INFORMATION FOR SEQ ID NO:349 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids(B) TYPE: amino acid 30 (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO:349: 35 Gln Gln Leu Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly His Glu 40 Lys Lys Gly Ile Ser Ser Thr Ser Glu Ser (350) INFORMATION FOR SEQ ID NO:350 45 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO:350:

5	of the Gin Ala Giu Leu Lys Pro Gly Lys Gly Lys Pro Glu Gln Glu  1 5 10 15
	Lys Lys Gly Thr Ser Ser Thr Ser Glu Ser 20 25
10	(351) INFORMATION FOR SEQ ID NO:351
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:351:
20	Gln Gln Gln Pro Glu Leu Lys Pro Gly Lys Gly Arg Asn Gly Lys Glu 1 5 10 15
25	Asn Lys Gly Lys Ser Ser Thr Ser Glu Ser 20 25
	(352) INFORMATION FOR SEQ ID NO:352
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:352:
	Gln Gln Gln Thr Glu Leu Arg Pro Gly Arg Gly Thr Thr Gly Gln Glu 1 5 10 15
40	Arg Lys Gly Lys Ser Ser Thr Ser Glu Ser
	(353) INFORMATION FOR SEQ ID NO:353
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:353:

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5	Gin His Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly His Glu 1 5 10 15
	Asn Lys Val Thr Ser Ser Thr Ser Glu Ser 20 25
10	(354) INFORMATION FOR SEQ ID NO:354
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
73	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:354:
20	Glu Gln Gln Ala Glu Leu Arg Ala Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
25	Gln Lys Ala Lys Ser Ser Thr Ser Glu Ser 20 25
20	(355) INFORMATION FOR SEQ ID NO:355
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:355:
	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
40	Lys Thr Gly Thr Ser Ser Thr Thr Glu Ser
	(356) INFORMATION FOR SEQ ID NO:356
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:356:

5	Gln Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Asn Pro Gly Gln Glu 1 5 10 15
	Lys Lys Ser Thr Ser Ser Ala Ser Glu Ser 20 25
10	(357) INFORMATION FOR SEQ ID NO:357
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:357:
20	Glu Gln Gln Thr Val Leu Arg Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
25	Lys Lys Gly Thr Ser Ala Thr Asn Glu Ser 20 25
	(358) INFORMATION FOR SEQ ID NO:358
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:358:
	Gln Gln Leu Thr Glu Leu Lys Pro Gly Asn Gly Thr Pro Gly Gln Glu 1 5 10 15
40	Lys Lys Ser Lys Ser Ser Thr Ser Glu Ser
	(359) INFORMATION FOR SEQ ID NO:359
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:359:

5	1 5 10 11 15 15 16 16 16 16 16 16 16 16 16 16 16 16 16
	Lys Lys Gly Thr Ser Ser Thr Ser Lys Ser 20 25
10	(360) INFORMATION FOR SEQ ID NO:360
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:360:
20	Leu Gln Gln Pro Val Leu Lys Pro Gly Lys Gly Ser His Gly Lys Gln 1 5 10 15
25	Lys Lys Asp Lys Ser Ser Thr Ser Glu Ser 20 25
	(361) INFORMATION FOR SEQ ID NO:361
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:361:
	Glu Gln Gln Pro Glu Thr Lys Pro Gly Lys Gly Thr Leu Gly Lys Gln 1 10 15
40	Lys Lys Ser Lys Ser Ser Thr Ser Glu Ser 20
	(362) INFORMATION FOR SEQ ID NO:362
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
Ju	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:362:

5	Gln Gln Gln Ala Glu Leu Lys Pro Gly Gln Gly Thr Pro Gly Gln Glu 1 10 15
J	Lys Lys Asn Lys Ser Ser Thr Pro Glu Phe 20 25
10	(363) INFORMATION FOR SEQ ID NO:363
4.5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:363:
20	Glu Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Asn Pro Glu Gln Pro 1 5 10 15
25	Lys Gln Gly Thr Ser Ser Thr Ser Glu Thr 20 25
	(364) INFORMATION FOR SEQ ID NO:364
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 364:
	Glu Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Asn Pro Glu Gln Pro 1 10 15
40	Lys Gln Gly Thr Ser Thr Thr Ser Glu Thr 20 25
	(365) INFORMATION FOR SEQ ID NO:365
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 365:

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5	Glu Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Asn Pro Glu Gln Pro 1 5 10 15
	Lys Gln Gly Thr Ser Ser Thr Ser Glu Thr 20 25
10	(366) INFORMATION FOR SEQ ID NO:366
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
,,	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:366:
20	Glu Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Asn Pro Glu Gln Pro 1 5 10 15
25	Lys Gln Asp Thr Ser Ser Thr Ser Glu Thr 20 25
	(367) INFORMATION FOR SEQ ID NO:367
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:367:
	Glu Gln Gln Ala Glu Leu Lys Pro Gly Lys Gly Asn Pro Glu Gln Pro 1 10 15
40	Lys Gln Gly Thr Ser Ser Thr Ser Gly Thr
	(368) INFORMATION FOR SEQ ID NO:368
<b>4</b> 5	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:368:

5	Glu Gln Gln Ala Glu Val Lys Pro Gly Lys Gly Asn Pro Glu Gln Pro 1 5 10 15
	Lys Gln Gly Thr Ser Ser Thr Ser Glu Thr 20 25
10	(369) INFORMATION FOR SEQ ID NO:369
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:369:
20	Glu Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Asn Pro Glu Gln Pro 1 5 10 15
25	Lys Gln Val Thr Ser Ser Thr Ser Glu Thr 20 25
	(370) INFORMATION FOR SEQ ID NO:370
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:370:
	Glu Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Asn Pro Glu Gln Pro 1 5 10 15
40	Lys Gln Ile Thr Ser Ser Thr Ser Glu Thr 20 25
	(371) INFORMATION FOR SEQ ID NO:371
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:371:

5	1 5 10 15
	Lys Gln Val Thr Ser Ser Thr Ser Glu Thr 20 25
10	(372) INFORMATION FOR SEQ ID NO:372
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:372:
20	Glu Gln Gln Ala Glu Leu Arg Pro Gly Arg Gly Asn Pro Glu Gln Pro 1 5 10 15
25	Lys His Val Thr Ser Ser Thr Ser Glu Thr 20 25
	(373) INFORMATION FOR SEQ ID NO:373
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:373:
	Glu Gln Gln Ala Glu Leu Arg Pro Gly Lys Gly Asn Thr Glu Gln Pro 1 5 10 15
40	Lys Gln Val Thr Ser Ser Thr Ser Glu Thr
	(374) INFORMATION FOR SEQ ID NO:374
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:374:

5	1 5 10 11 11 11 11 11 11 11 11 11 11 11 11
	Lys Leu Ile Thr Ser Ser Thr Ser Glu Thr 20 25
10	(375) INFORMATION FOR SEQ ID NO:375
15	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:375:
20	Thr Gly Gln Ala Glu Leu Arg Pro Gly Lys Gly Ala Pro Glu Gln Gly 1 5 10 15
25	Lys Lys Gly Lys Ser Ser Thr Ser Asp Arg 20 25
	(376) INFORMATION FOR SEQ ID NO:376
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:376:
	Gln Tyr Gln Ala Glu Leu Arg Pro Gly Lys Gly Thr Pro Arg Gln Gln 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser
	(377) INFORMATION FOR SEQ ID NO:377
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:377:

	Gln Gln Gln Ala Val Leu Arg His Gly Lys Gly Thr His Gly Gln Glu 1 5 10 15
5	Lys Lys Gly Lys Ser Ser Thr Ser Glu Ser
	20 25
10	(378) INFORMATION FOR SEQ ID NO:378
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
13	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:378:
20	Gln Gln Gln Thr Lys Leu Gly Pro Gly Arg Gly Thr Pro Gly Gln Gly 1 5 10 15
25	Arg Lys Gly Lys Ser Ser Thr Ser Gly Ser 20 25
	(379) INFORMATION FOR SEQ ID NO:379
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:379:
	Glu Gln Gln Ala Glu Leu Arg Ala Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
40	Lys Lys Gly Lys Ser Ser Val Tyr Phe Ala 20 25
	(380) INFORMATION FOR SEQ ID NO:380
<b>4</b> 5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:380:

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5	Glu Gln Gln Ala Glu Leu Lys Ala Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
	Lys Gln Gly Glu Ser Thr Arg Ser Glu Thr 20 25
10	(381) INFORMATION FOR SEQ ID NO:381
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:381:
20	Gln Gln Lys Ala Glu Leu Ala Ala Ser Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
25	Lys Lys Gly Arg Ser Ser Thr Ser Glu Ser 20 25
	(382) INFORMATION FOR SEQ ID NO:382
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:382:  Gln Gln Gln Thr Glu Leu Arg Pro Gly Lys Gly Thr Pro Gly Gln Glu
	1 5 10 15 15
40	Lys Arg Gly Lys Ser Ser Asn Leu Arg Leu 20 25
	(383) INFORMATION FOR SEQ ID NO:383
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:383:

5	Glu Lys Val Gly Gly Leu Gln Gly Ser Ser Phe Asp Pro Gly Lys Ala 1 5 10 15
	Ser Lys Gly Thr Ser Gln Arg Ala Glu Thr 20 25
10	(384) INFORMATION FOR SEQ ID NO:384
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
7.5	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:384:
20	Glu Gln Gln Ala Asp Leu Lys Leu Gly Lys Gly Asn Pro Glu Gln Pro 1 5 10 15
25	Lys Leu Ala Thr Pro Ser Thr Ser Glu Thr 20 25
	(385) INFORMATION FOR SEQ ID NO:385
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:385:
٠	Glu Gln Val Gly Gly Leu Lys Pro Gly Lys Gly Thr Pro Asp Lys Ser 1 5 10 15
40	Asp Val Lys Asp Asn Ala Lys Ser Glu Thr
	(386) INFORMATION FOR SEQ ID NO:386
<b>4</b> 5	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
•	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:386:

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5	Asp Gln Gln Pro Asp Leu Lys Pro Ser Ser Gly Ser Pro Gly His Pro 1 5 10 15
	Ser Lys Ser Thr Ser Lys Thr Thr Glu Thr 20 25
10	(387) INFORMATION FOR SEQ ID NO:387
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:387:
20	Asp Gln Gln Pro Asp Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
25	Ser Lys Ser Thr Ser Lys Thr Thr Glu Thr 20 25
	(388) INFORMATION FOR SEQ ID NO:388
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:388:
	Asp Gln Gln Pro Asp Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
40	Ser Lys Ser Thr Ser Lys Thr Ala Glu Thr 20 25
	(389) INFORMATION FOR SEQ ID NO:389
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:389:

5	Asp Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
	Ser Lys Ser Thr Ser Lys Thr Thr Glu Thr 20 25
10	(390) INFORMATION FOR SEQ ID NO:390
15	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:390:
20	Asp Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
25	Ser Lys Asn Thr Ser Lys Thr Thr Glu Thr 20 25
	(391) INFORMATION FOR SEQ ID NO:391
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:391:
	Asp Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asp Pro 1 5 10 15
40	Ser Lys Thr Thr Ser Lys Thr Thr Glu Thr
	(392) INFORMATION FOR SEQ ID NO:392
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
<b>5</b> 0	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:392:

5	Asp Gin Gin Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro  1 10 15
	Ser Lys Thr Thr Ser Lys Thr Thr Glu Thr 20 25
10	(393) INFORMATION FOR SEQ ID NO:393
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:393:
20	Asp His Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 10 15
25	Ser Lys Asn Thr Ser Lys Thr Thr Glu Thr 20 25
	(394) INFORMATION FOR SEQ ID NO:394
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:394:
	Asp Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
<b>40</b>	Ser Arg Ser Thr Ser Lys Thr Thr Glu Thr
	(395) INFORMATION FOR SEQ ID NO:395
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:395:

	Asp Gln Gln Pro Gly Leu Lys Pro Ser Ala Gly Ser Pro Gly Asn Pro 1 5 10 15
5	Con Ites Con Whee Con The Control of the Control of
	Ser Lys Ser Thr Ser Lys Thr Ala Glu Thr 20 25
10	(396) INFORMATION FOR SEQ ID NO:396
	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:396:
20	Glu Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
25	Ser Lys Ser Thr Ser Lys Thr Ser Glu Thr 20 25
	(397) INFORMATION FOR SEQ ID NO:397
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:397:
	Asp Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
40	Ser Lys Asn Thr Ser Lys Thr Ile Glu Thr 20 25
	(398) INFORMATION FOR SEQ ID NO:398
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:398:

5	Asp Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asp Pro 1 5 10 15
	Ser Lys Asn Thr Ser Lys Thr Pro Glu Thr
10	(399) INFORMATION FOR SEQ ID NO:399
15	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:399:
20	Glu Gln Gln Pro Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
25	Ser Lys Ser Thr Ser Lys Thr Thr Glu Thr 20 25
	(400) INFORMATION FOR SEQ ID NO:400
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:400:
	Asp Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
40	Ser Lys Asn Thr Ser Glu Thr Thr Glu Thr
	(401) INFORMATION FOR SEQ ID NO:401
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:401:

5	Asp Gin Gin Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro  1 5 10 15
	Ser Lys Asn Thr Ser Glu Thr Thr Glx Thr 20 25
10	(402) INFORMATION FOR SEQ ID NO:402
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 402:
20	Glu Gln Gln Pro Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
25	Ser Lys Ser Thr Ser Lys Thr Ser Glu Thr 20 25
	(403) INFORMATION FOR SEQ ID NO:403
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:403:
	Glu Gln Gln Pro Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
<b>1</b> 0	Ser Lys Ser Thr Ser Arg Thr Thr Glu Thr
	(404) INFORMATION FOR SEQ ID NO:404
<b>1</b> 5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:404:

5	1 5 10 Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro
	Ser Lys Ser Thr Ser Lys Thr Ala Glu Thr 20 25
10	(405) INFORMATION FOR SEQ ID NO: 405
15	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:405:
20	Asp Gln Gln Pro Asp Leu Lys Pro Ser Ser Gly Phe Pro Gly Asn Pro 1 5 10 15
25	Ser Lys Ser Thr Ser Lys Thr Thr Glu Thr 20 25
	(406) INFORMATION FOR SEQ ID NO:406
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:406:
	Glu Gln Gln Pro Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly Lys Pro 1 5 10 15
40	Ser Lys Ser Thr Ser Lys Thr Asn Glu Thr
	(407) INFORMATION FOR SEQ ID NO:407
<b>4</b> 5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:407:

5	Glu Gln Gln Pro Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
10	Ser Lys Ser Thr Phe Lys Thr Ser Glu Thr 20 25
10	(408) INFORMATION FOR SEQ ID NO:408
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:408:
20	Glu Gln Gln Pro Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
25	Ser Lys Ser Thr Ser Thr Thr Ser Glu Thr 20 25
	(409) INFORMATION FOR SEQ ID NO:409
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(X1) SEQUENCE DESCRIPTION: SEQ ID NO:409:
	Glu Gln Gln Leu Ser Leu Lys Pro Ser Ser Gly Ser Pro Gly Asn Pro 1 5 10 15
40	Ser Lys Ser Thr Ser Lys Thr Thr Glu Thr
	(410) INFORMATION FOR SEQ ID NO:410
<b>45</b>	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:410:

5	Gln Gln Gln Pro Gly Leu Lys Pro Ser Phe Gly Pro Pro Gly Lys Pro 1 5 10 15
	Ser Gln Ser Thr Ser Lys Thr Thr Glu Thr 20 25
10	(411) INFORMATION FOR SEQ ID NO:411
	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:411:
20	Gln Gln Lys Pro Gly Leu Ala Pro Ser Ser Gly Ser Pro Gly Lys Ser 1 5 10 15
	Thr Lys Ser Asn Ser Lys Gln Thr Asp Thr 20 25
25	(412) INFORMATION FOR SEQ ID NO:412
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:412:
	Gln Gln Lys Pro Gly Leu Ala Pro Ser Ser Gly Ser Pro Gly Lys Ser 1 10 15
40	Ala Lys Ser Asn Ser Lys Gln Thr Asp Thr 20 25
	(413) INFORMATION FOR SEQ ID NO:413
<b>45</b> .	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:413:
	Gln Gln Lys Pro Gly Leu Ala Pro Ser Ser Gly Ser Pro Gly Lys Ser 1 10 15
55	

5	Ala Met Ser Asn Ser Lys Gln Thr Asp Thr 20 25
	(414) INFORMATION FOR SEQ ID NO:414
10	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:414:
	Gln Gln Lys Pro Gly Leu Ala Pro Ser Ser Gly Ser Pro Gly Lys Ser 1 5 10 15
20	Ala Ile Ser Asn Ser Lys Gln Thr Asp Thr 20 25
	(415) INFORMATION FOR SEQ ID NO:415
25	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
30	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:415:
05	Gln Gln Lys Pro Gly Leu Gln Pro Ser Ser Gly Ser Pro Gly Lys Ala 1 5 10 15
35	Ala Ile Ser Asn Ser Lys Gln Ser Asn Thr 20 25
	(416) INFORMATION FOR SEQ ID NO:416
40	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:416:
50	Gln Gln Lys Pro Gly Leu Gln Pro Ser Ser Gly Ser Pro Gly Lys Ala 1 5 10 15
	Ala Il Ser Asn Ser Lys Gln Ala Asn Thr 20 25
55	

	(417) INFORMATION FOR SEQ ID NO:417
5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:417:
15	Gln Gln Lys Pro Val Leu Ala Pro Ser Ser Gly Ser Pro Gly Lys Ser 1 5 10 15
	Ala Met Ser Asn Ser Lys Gln Ile Asp Thr 20 25
20	(418) INFORMATION FOR SEQ ID NO:418
25	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:418:
30	Gln Gln Lys Pro Ser Leu Gln Pro Ser Ser Asp Ser Pro Gly Lys Ala 1 5 10 15
<i>30 35</i>	
	1 5 10 15  Ala Met Ser Asn Ser Lys Gln Ala Asp Thr
	1 5 10 15  Ala Met Ser Asn Ser Lys Gln Ala Asp Thr 20 25
35	Ala Met Ser Asn Ser Lys Gln Ala Asp Thr 20 25  (419) INFORMATION FOR SEQ ID NO:419  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid
35	Ala Met Ser Asn Ser Lys Gln Ala Asp Thr 20 25  (419) INFORMATION FOR SEQ ID NO:419  (i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear
<b>35</b>	Ala Met Ser Asn Ser Lys Gln Ala Asp Thr 20 25  (419) INFORMATION FOR SEQ ID NO:419  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear  (ii) MOLECULE TYPE: peptide
<b>35</b>	Ala Met Ser Asn Ser Lys Gln Ala Asp Thr 20 25  (419) INFORMATION FOR SEQ ID NO:419  (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid (C) TOPOLOGY: linear  (ii) MOLECULE TYPE: peptide  (xi) SEQUENCE DESCRIPTION: SEQ ID NO:419:  Glu Arg Val Gly Asp Leu Glu Pro Gly Arg Gly Ile Pro Gly Lys Ala

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5	(1) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: peptide	
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 420:	
	Glu Arg Val Gly Asp Leu Glu Pro Glu Arg Gly Ile Pro Gly Ly 1 5 10	s Ala .15
15	Pro Lys Gly Asp Ser Lys Lys Ile Glu Thr 20 25	
	(421) INFORMATION FOR SEQ ID NO:421	
20	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>	
25	(ii) MOLECULE TYPE: peptide	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:421:	
30	Glu Gln Val Gly Gly Leu Lys Pro Gly Arg Gly Thr Pro Gly Ly 1 10	s Ala 15
	Pro Lys Gly Asp Ser Lys Lys Thr Glu Thr 20 25	
35	(422) INFORMATION FOR SEQ ID NO: 422	
	<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 26 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul>	••
40	(ii) MOLECULE TYPE: peptide	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:422:	
45	Glu Gln Val Gly Gly Leu Gln Pro Gly Lys Gly Thr Ser Gly Ly 1 5 10	s Ala 15
50	Ser Lys Gly Asp Ser Lys Lys Thr Glu Thr 20 25	
	(423) INFORMATION FOR SEQ ID NO:423	·
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids	
55	(11) DENOTIFE 20 GHITHO GCIUS	

	(B) TYPE: amino acid (C) TOPOLOGY: linear
5	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 423:
10	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asx 1 5 10 15
15	Ser Lys Gly Asp Ser Lys Arg Ala Glu Thr 20 25
	(424) INFORMATION FOR SEQ ID NO:424
20	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:424:
	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
30	Ser Lys Gly Asn Ser Lys Arg Ala Glu Thr 20 25
	(425) INFORMATION FOR SEQ ID NO:425
35	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
40	(ii) MOLECULE TYPE: peptide
	(Hi) SEQUENCE DESCRIPTION: SEQ ID NO: 425:
45	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
	Ser Arg Gly Asn Ser Lys Arg Ala Glu Thr 20 25
50	(426) INFORMATION FOR SEQ ID NO:426
	<ul> <li>(i) SEQUENCE CHARACTERISTICS:         <ul> <li>(A) LENGTH: 26 amino acids</li> <li>(B) TYPE: amino acid</li> <li>(C) TOPOLOGY: linear</li> </ul> </li> </ul>
55	(C) IOFOLOGI. IINEGI

_	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 426:
10	Glu Gln Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
,,,	Ser Lys Gly Asn Ser Lys Arg Ala Glu Thr 20 25
	(427) INFORMATION FOR SEQ ID NO:427
15	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:427:
25	Glu Gln Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
	Ser Lys Gly Asn Ala Lys Arg Ala Glu Thr 20 25
30	(428) INFORMATION FOR SEQ ID NO:428
35	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 428:
<b>4</b> 0	Glu Gln Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp  1 10 15
45	Ser Lys Gly Asp Ser Arg Arg Ala Glu Thr 20 25
	(429) INFORMATION FOR SEQ ID NO:429
50	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide

	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 429:
<b>5</b>	Glu Gln Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
10	Ser Lys Gly Asn Ser Arg Arg Ala Glu Thr 20 25
	(430) INFORMATION FOR SEQ ID NO:430
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:430:
	Gln Gln Val Gly Gly Leu Glu Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
25	Ser Lys Gly Ask Ser Lys Arg Ala Glu Thr 20 25
	(431) INFORMATION FOR SEQ ID NO:431
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:431:
40	Glu Gln Leu Gly Asp Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Ala 1 5 10 15
	Ser Lys Glý Asn Ser Lys Arg Ala Glu Thr 20 25
45	(432) INFORMATION FOR SEQ ID NO:432
50	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:432:
55	

5.	Glu Gln Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Thr Gly Lys Asp 1 5 10 15
	Ser Lys Gly Asp Ser Lys Arg Ala Glu Thr 20 25
10	(433) INFORMATION FOR SEQ ID NO:433
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:433:
20	Gln Gln Val Gly Gly Val Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
25	Ser Lys Gly Asn Ser Lys Arg Ala Glu Thr 20 25
	(434) INFORMATION FOR SEQ ID NO:434
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
`	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:434:
	Gln Gln Val Gly Gly Val Gln Pro Gly Arg Gly Ile Pro Gly Lys Asp 1 5 10 15
40	Ser Lys Gly Asn Ser Lys Arg Pro Glu Thr
	(435) INFORMATION FOR SEQ ID NO:435
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:435:

. 5	Glu Gln Val Gly Gly Val Gln Pro Gly Arg Gly Ile Pro Gly Lys Asp 1 5 10 15
	Ser Lys Gly Asp Ser Lys Arg Pro Glu Thr 20 25
10	(436) INFORMATION FOR SEQ ID NO:436
	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:436:
20	Gln Gln Val Gly Gly Val Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
	Ser Asn Gly Asp Ser Lys Arg Pro Glu Thr 20 25
25	(437) INFORMATION FOR SEQ ID NO:437
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:437:
30	Gln Lys Val Gly Gly Val Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
40	Ser Lys Gly Asn Ser Lys Arg Thr Glu Thr 20 25
	(438) INFORMATION FOR SEQ ID NO:438
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:438:
	Gln Glu Val Gly Gly Val Glx Pro Gly Arg Gly Thr Pro Gly Lys Asx 1 5 10 15
55	

5	Ser Lys Gly Asx Ser Lys Arg Ala Glu Thr 20 25
	(439) INFORMATION FOR SEQ ID NO:439
10	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:439:
	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
20	Ser Asn Gly Asp Ser Lys Gln Ala Glx Thr 20 25
	(440) INFORMATION FOR SEQ ID NO:440
25	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
30	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:440:
35	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Ser Pro Gly Lys Asp 1 5 10 15
	Thr Asn Gly Asp Ser Lys Glu Ala Glx Thr 20 25
40	(441) INFORMATION FOR SEQ ID NO:441
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:441:
50	Ala Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 10 15
i5	Ser Asn Gly Asp Ser Lys Gln Ala Glx Ser 20 25

	(442) INFORMATION FOR SEQ ID NO:442
5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:442:
15	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Va. 1 5 10 15
	Ser Gln Gly Asp Ser Lys Gln Ala Glx Thr 20 25
20	(443) INFORMATION FOR SEQ ID NO:443
25	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:443:
30	Glu Gln Val Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Val
35	Ser Gln Gly Asp Ser Lys Glu Pro Glx Thr 20 25
	(444) INFORMATION FOR SEQ ID NO: 444
40	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
45	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:444:
	Glu Gln Leu Gly Gly Leu Gln Pro Glu Arg Gly Thr Pro Gly Lys Glu 1 5 15
50	Ser Lys Gly Asn Ser Met Arg Ala Glu Thr 20 25
	(445) INFORMATION FOR SEQ ID NO:445

5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:445:
	Glu Gln Val Gly Asp Leu Gln Pro Gly Arg Gly Asx Pro Gly Lys Asp 1 5 10 15
15	Ser Lys Gly Asn Ala Lys Arg Val Glu Thr 20 25
	(446) INFORMATION FOR SEQ ID NO:446
20	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
25	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:446:
30	Glu Gln Val Gly Asp Leu Gln Pro Gly Arg Gly Asn Pro Gly Lys Asp 1 5 10 15
	Ser Lys Gly Asn Ala Gln Arg Pro Glu Thr 20 25
35	(447) INFORMATION FOR SEQ ID NO:447
40	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:447:
<b>4</b> 5	Gln Gln Val Gly Gly Val Gln Pro Gly Arg Gly Thr Leu Gly Lys Asp 1 5 10 15
50	Ser Lys Gly Asn Ser Lys Arg Ala Glu Thr 20 25
	(448) INFORMATION FOR SEQ ID NO:448
E.E	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids

_	(C) TOPOLOGY: linear
5	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:448:
10	Gln Glx Val Gly Gly Ala Glx Pro Gly Arg Gly Ser Pro Gly Lys Ala 1 5 10 15
15	Ser Lys Gly Asx Ser Lys Arg Ala Glu Thr 20 25
	(449) INFORMATION FOR SEQ ID NO:449
20	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:449:
	Gln Gln Val Gly Gly Leu Lys Pro Gly Arg Gly Ser Pro Gly Lys Asp 1 5 10 15
30	Ser Lys Gly Asn Ala Gln Arg Thr Glx Thr 20 25
	(450) INFORMATION FOR SEQ ID NO:450
35	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
40	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 450:
<b>4</b> 5	Asp Gln Val Gly Gly Leu Lys Pro Gly Arg Gly Thr Pro Gly Lys Asn 1 5 10 15
	Ser Asn Gly Asp Ser Lys Thr Pro Glx Thr 20 25
50	(451) INFORMATION FOR SEQ ID NO: 451
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids (B) TYPE: amino acid
55	(C) TOPOLOGY: linear

	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:451:
	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Ser Arg Glu Asp 1 5 10 15
10	Ser Lys Gly Asn Ser Lys Arg Ala Glu Thr 20 25
	(452) INFORMATION FOR SEQ ID NO:452
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
20	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:452:
25	Glu Gln Val Gly Ala Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Asp 1 5 10 15
	Ser Gln Ala Asp Ser Lys Glu Ala Glx Thr 20 25
30	(453) INFORMATION FOR SEQ ID NO:453
35	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 22 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:453:
40	Glu Gln Leu Gly Gly Leu Gln Pro Gly Arg Gly Thr Pro Gly Lys Val 1 _ 5 10 15
<b>4</b> 5	Glu Gly Ser Val Glu Thr 20
	(454) INFORMATION FOR SEQ ID NO:454
50	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
55	

	(x1) SEQUENCE DESCRIPTION: SEQ ID NO:454:
5	Glu Gln Val Gly Ala Phe Gln Pro Gly Arg Gly Asn Ser Gly Lys Ala 1 5 10 15
10	Ser Lys Gly Asp Ser Lys Arg Pro Asp Thr 20 25
	(455) INFORMATION FOR SEQ ID NO:455
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 455:
	Glu Gln Val Gly Ala Phe Gln Pro Gly Lys Gly Asn Ser Gly Lys Ala 1 5 10 15
25	Ser Lys Gly Asp Ser Lys Arg Pro Asp Thr 20 25
	(456) INFORMATION FOR SEQ ID NO:456
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 456:
	Glu Gln Val Gly Ala Phe Gln Pro Gly Lys Gly Asn Ser Gly Lys Ala
40	1 5 10 15
	Ser Lys Gly Asp Ser Asn Arg Pro Asp Thr 20 25
45	(457) INFORMATION FOR SEQ ID NO:457
50	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:457:
55	

5	Gln Gln Val Gly Gly Val Gln Ala Gly Arg Ala Asn Pro Gly Lys Asp 1 5 10 15
	Ser Arg Gly Ile Ser Lys Arg Thr Glu Thr 20 25
10	(458) INFORMATION FOR SEQ ID NO:458
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:458:
20	Gln Gln Val Ala Glu Val Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
25	Lys Gln Gly Glu Ser Thr Arg Ser Glu Thr 20 25  (459) INFORMATION FOR SEQ ID NO:459
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
25	<ul><li>(ii) MOLECULE TYPE: peptide</li><li>(xi) SEQUENCE DESCRIPTION: SEQ ID NO:459:</li></ul>
35	Gln Gln Val Ala Glu Val Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
40	Lys Gln Gly Thr Ser Thr Arg Ser Glu Thr
	(460) INFORMATION FOR SEQ ID NO:460
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide  (xi) SEQUENCE DESCRIPTION: SEQ ID NO:460:

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5	Gln Gln Val Ala Glu Val Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
	Lys Gln Gly Thr Ser Ala Arg Ser Glu Thr 20 25
	(461) INFORMATION FOR SEQ ID NO:461
10	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
15	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:461:
20	Gln Gln Val Ala Glu Val Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
	Lys Gln Gly Thr Ser Ile Arg Ser Asp Thr 20 25
25	(462) INFORMATION FOR SEQ ID NO:462
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:462:
35	Gln Gln Val Ala Glu Val Lys Pro Gly Lys Gly Thr Pro Gly Gln Glu 1 5 10 15
<b>4</b> 0	Lys Gln Gly Thr Ser Ile Arg Ser Asp Thr 20 25
	(463) INFORMATION FOR SEQ ID NO:463
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:463:
	Gln Gln Val Ala Glu Val Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
55	

	Asn Gln Gly Thr Ser Thr Arg Ser Asp Thr 20 25
5	(464) INFORMATION FOR SEQ ID NO:464
10	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 464:
15	Gln Gln Val Gly Glu Val Lys Pro Gly Arg Gly Thr Pro Gly Gln Gln 1 5 10 15
20	Lys Gln Asp Thr Ser Thr Arg Ser Asp Thr 20 25
	(465) INFORMATION FOR SEQ ID NO:465
25	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:465:
	Gln Gln Val Ala Glu Val Lys Pro Gly Arg Gly Thr Pro Gly His Pro 1 5 10 15
35	Arg Gln Gly Ala Ser Phe Arg Ser Asp Ser 20 25
	(466) INFORMATION FOR SEQ ID NO:466
40	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:466:
50	Gln Gln Val Ser Glu Leu Lys Pro Gly Lys Gly Thr Pro Gly Gln Gln 1 5 10 15
	Gly Thr Gly Thr Ser Val Lys Ala Glu Thr 20 25
EE	

	(467) INFORMATION FOR SEQ ID NO:467
5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:467:
15	Glu Gln Val Ala Glu Val Lys Pro Gly Lys Gly Ser Pro Gly Lys Pro 1 5 10 15
	Ser Gln Gly Lys Ser Ile Lys Ala Ser Thr 20 25
20	(468) INFORMATION FOR SEQ ID NO:468
25	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:468:
30	Glu Gln Val Ala Glu Val Lys Pro Gly Arg Gly Ser Pro Gly Lys Pro 1 5 10 15
35	Ser Gln Gly Lys Ser Ile Lys Ala Ser Thr 20 25
	(469) INFORMATION FOR SEQ ID NO: 469
40	<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 26 amino acids</li> <li>(B) TYPE: amino acid</li> <li>_ (C) TOPOLOGY: linear</li> </ul>
	(ii) MOLECULE TYPE: peptide
45	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:469:
	Gln Gln Val Ala Glu Val Lys Pro Gly Arg Gly Asp Pro Gly Arg Pro 1 5 10 15
50	Arg Gln Ala Ser Ser Thr Ile Ser Ala Thr 20 25
	(470) INFORMATION FOR SEQ ID NO:470

5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 470:
	Glu Gln Val Ala Glu Val Pro Gln Gly Lys Gly Arg Pro Gly Lys Ser 1 5 10 15
15	Leu Gln Gly Lys Ser Leu Lys Ala Ser Thr 20 25
	(471) INFORMATION FOR SEQ ID NO:471
20	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
25	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:471:
30	Gln Gln Met Ala Glu Val Lys Pro Gly Arg Gly Thr Pro Gly Lys Pro 1 5 10 15
	Gly Val Val Pro Ser Phe Phe Ser Glu Thr 20 25
35	(472) INFORMATION FOR SEQ ID NO:472
40	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
40	( <u>i</u> i) MOLECULE TYPE: peptide
-	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 472:
<b>4</b> 5	Gln Gln Val Ala Glu Val Lys Pro Gly Arg Gly Thr Pro Gly Arg Tyr 1 5 10 15
50	Ile Trp Glu Pro Ser Phe Phe Asn Glu Gly 20 25
	(473) INFORMATION FOR SEQ ID NO:473
55	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 amino acids

5	(B) TYPE: amino acid (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:473:
10	Gln Gln Gln Ala Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Lys Pro 1 5 10 15
15	Ser Lys Ser Thr Ser Lys Thr Ala Ala Thr 20 25
	(474) INFORMATION FOR SEQ ID NO:474
20	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:474:
	Gln Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Lys Pro 1 5 10 15
30	Ser Lys Ser Thr Ser Lys Thr Ala Ala Thr 20 25
	(475) INFORMATION FOR SEQ ID NO:475
35	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
40	(#i)_SEQUENCE DESCRIPTION: SEQ ID NO:475:
	Gln Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Lys Pro 1 5 10 15
45	Ser Lys Ser Thr Ser Asn Thr Ala Ala Thr 20 25
	(476) INFORMATION FOR SEQ ID NO:476
50	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	·

	(ii) MOLECULE TYPE: peptide	
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:476:	
	Gln Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Ala Gly Lys Pr 1 5 15	0
10	Ser Lys Ser Thr Ser Lys Thr Ala Ala Thr 20 25	
	(477) INFORMATION FOR SEQ ID NO:477	
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>	
20	(ii) MOLECULE TYPE: peptide	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:477:	
25	Arg Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Pro Pro Gly Lys Pro 1 5 10 15	0
	Ser Arg Gly Thr Ser Arg Ser Ala Ala Thr 20 25	
30	(478) INFORMATION FOR SEQ ID NO:478	
35	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>	
	(ii) MOLECULE TYPE: peptide	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:478:	
40	Gln Gln Gln Ala Gly Leu Lys Pro Ser Ser Gly Ser Pro Gly Arg Thi	r.
45	Ser Lys Ser Thr Ser Lys Thr Ala Ala Thr 20 25	
	(479) INFORMATION FOR SEQ ID NO:479	
50	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>	
55	(ii) MOLECULE TYPE: peptide	

	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:479:
5	Gln Gln Glu Pro Gly Leu Arg Pro Ser Ser Gly Thr Pro Gly Arg Thr 1 5 10 15
10	Pro Arg Ser Thr Ser Lys Thr Ala Ala Thr 20 25
	(480) INFORMATION FOR SEQ ID NO:480
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:480:
	Xaa Gln Glu Pro Gly Leu Arg Pro Ser Ser Gly Ser Pro Gly Arg Thr 1 5 10 15
25	
	Pro Arg Ser Thr Ser Lys Thr Ala Ala Thr 20 25
	(481) INFORMATION FOR SEQ ID NO:481
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:481:
	Gln Gln Gln Pro Gly Leu Lys Pro Ser Ser Gly Ser Pro Ser Arg Val
40	1 5 10 15
	Ser Lys Ser Thr Ser Lys Thr Pro Glu Thr 20 25
45	(482) INFORMATION FOR SEQ ID NO:482
50	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:482:
55	

<b>5</b> .	Gln His Gln Ala Gly Leu Lys Arg Ser Ser Gly Pro Pro Gly Lys Pr 1 10 15	0
	Ser Thr Ser Thr Ser Lys Thr Ala Ala Thr 20 25	
10	(483) INFORMATION FOR SEQ ID NO:483	
<b>1</b> 5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>	
	(ii) MOLECULE TYPE: peptide	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:483:	
20	Glx Gln Glu Ser Gly Leu Lys Pro Thr Ser Gly Ser Pro Gly Lys Pro 1 5 10 15	0
25	Ser Lys Ser Arg Ser Lys Ala Ala Asp Ala 20 25	
	(484) INFORMATION FOR SEQ ID NO:484	
30	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>	
	(ii) MOLECULE TYPE: peptide	
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:484:	
and the	Gln Thr Lys Pro Thr Leu Lys Pro Thr Thr Gly Ser Pro Gly Arg Pro 1 5 10 15	5
40	Ser Lys Ser Thr Ser Lys Asp Pro Val Thr	
	(485) INFORMATION FOR SEQ ID NO:485	
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear	
50	(ii) MOLECULE TYPE: peptide	
-	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:485:	

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5	Gln Thr Lys Pro Thr Leu Lys Pro Thr Thr Gly Ser Pro Gly Lys Pro 1 5 10 15
	Ser Arg Ser Thr Ser Arg Asp Pro Val Ser 20 25
10	(486) INFORMATION FOR SEQ ID NO:486
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
,,,	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:486:
20	Glu Thr Arg Pro Ala Leu Lys Pro Thr Thr Gly Ser Pro Gly Lys Thr 1 5 10 15
	Ser Lys Thr Thr Ser Lys Asp Pro Val Thr
25	(487) INFORMATION FOR SEQ ID NO:487
30	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:487:
35	Gln Asn Arg Pro Ala Leu Lys Ala Thr Thr Gly Ser Pro Gly Lys Thr 1 5 10 15
40	Ser Glu Thr Thr Ser Lys Asp Pro Ala Thr 20 25
	(488) INFORMATION FOR SEQ ID NO: 488
45	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:488:
	Gln Thr Thr Pro Ala Leu Lys Pro Lys Thr Gly Ser Pro Gly Lys Thr 1 5 10 15

5	Ser Arg Thr Asp Ser Lys Asn Pro Val Thr 20 25
	(489) INFORMATION FOR SEQ ID NO:489
10	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:489:
	Gln Thr Arg Pro Ala Leu Arg Pro Thr Thr Gly Ser Pro Gly Glu Ala 1 5 10 15
20	Ser Glu Thr Thr Ser Lys Gly Pro Gly Thr 20 25
	(490) INFORMATION FOR SEQ ID NO:490
25	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 26 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 490:
	Gln Thr Arg Pro Ala Leu Lys Pro Thr Thr Gly Ser Pro Gly Lys Thr 1 5 10 15
35	Ser Glu Thr Thr Ser Arg Asp Thr Ala Tyr 20 25
	(491) INFORMATION FOR SEQ ID NO:491
40	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 26 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
45	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:491:
50	Leu Glu Gly Val Gln Leu Trp Gly Gly Arg Gly Ile Ser Arg Lys Tyr 1 5 10 15
	Ala Lys Gly Asn Gly Lys Arg Glu Asp Ser 20 25
55	

	(492) INFORMATION FOR SEQ ID NO:492
5	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 10 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:492:
	Tyr Asn Asn Pro Gly Asn Gly Tyr Ile Ala 1 5 10
15	(493) INFORMATION FOR SEQ ID NO:493
20	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 10 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:493:
	Tyr Ile Asn Pro Gly Lys Gly Tyr Leu Ser 1 5 10
30	(494) INFORMATION FOR SEQ ID NO:494
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 11 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
35	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:494:
40	Arg Ala Ser Gln Glu Ile Ser Gly Tyr Leu Ser 1 5 10
	(495) INFORMATION FOR SEQ ID NO:495
<b>4</b> 5	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 11 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:495:
55	Arg Ala Ser Gly Asn Ile His Asn Tyr Leu Ala 1 5 10

	(496) INFORMATION FOR SEQ ID NO:496
5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 11 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:496:
	Arg Ala Ser Gln Asp Ile Asn Asn Phe Leu Asn 1 5 10
15	(497) INFORMATION FOR SEQ ID NO:497
20	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 11 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:497:
	Arg Ala Ser Gln Ser Ile Gly Asn Asn Leu His 1 5 10
30	(498) INFORMATION FOR SEQ ID NO:498
35	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 7 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:498:
40	Ala Ala Ser Thr Leu Asp Ser 1 5
	(499) INFORMATION FOR SEQ ID NO:499
<b>4</b> 5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 7 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:499:
55	Tyr Thr Thr Leu Ala Asp

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	(500) INFORMATION FOR SEQ ID NO:500
5	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 7 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:500:
	Phe Thr Ser Arg Ser Gln Ser 1 5
15	(501) INFORMATION FOR SEQ ID NO:501
20	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 7 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:501:
	Lys Ala Ser Ser Leu Glu Ser 1 5
30	(502) INFORMATION FOR SEQ ID NO:502
. 35	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 9 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:502:
40	Leu Gln Tyr Leu Ser Tyr Pro Leu Thr 1 5
	(503) INFORMATION FOR SEQ ID NO:503
<b>4</b> 5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 9 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:503:
55	Gln His Phe Trp Ser Thr Pro Arg Thr

	(504) INFORMATION FOR SEQ ID NO:504
5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 9 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:504:
	Gln Gln Gly Asn Ala Leu Pro Arg Thr 1 5
15	(505) INFORMATION FOR SEQ ID NO:505
20	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 7 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:505:
	Gln Gln Tyr Asn Ser Tyr Ser 1 5
30	(506) INFORMATION FOR SEQ ID NO:506
35	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 5 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:506:
40	Thr Phe Gly Ile Thr 1 5
	(507) INFORMATION FOR SEQ ID NO:507
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 5 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:507:
55	Gly Tyr Gly Val Asn 5

	(508) INFORMATION FOR SEQ ID NO:508
5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 5 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:508:
15	Ser Asn Gly Ile Asn 1 5
	(509) INFORMATION FOR SEQ ID NO:509
20	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 5 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:509:
	Asp Tyr Ala Met His 1 5
30	(510) INFORMATION FOR SEQ ID NO:510
35	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 10 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:510:
40	Glu Ile Phe Pro Gly Asn Ser Lys Thr Tyr 1 5 10
	(511) INFORMATION FOR SEQ ID NO:511
45	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 9 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:511:
55	Met Ile Trp Gly Asp Gly Asn Thr Asp

	(512) INFORMATION FOR SEQ ID NO:512
5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 10 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
10 .	(ii) MOLECULE TYPE: peptide
70,	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:512:
	Tyr Asn Asn Pro Gly Asn Gly Tyr Ile Ala 1 5 10
15	(513) INFORMATION FOR SEQ ID NO:513
20	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 9 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:513:
	Ile Ser Trp Asp Ser Ser Ser Ile Gly 1 5
30	(514) INFORMATION FOR SEQ ID NO:514
35	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 5 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
 	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:514:
40	Arg Glu Ile Arg Tyr 1 5
	(515) INFORMATION FOR SEQ ID NO:515
<b>45</b>	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 8 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
50	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:515:
55	Glu Arg Asp Tyr Arg Leu Asp Tyr

	(516) INFORMATION FOR SEQ ID NO:516
5	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 12 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
10	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:516:
15	Ser Glu Tyr Tyr Gly Gly Ser Tyr Lys Phe Asp Tyr 1 5 10
	(517) INFORMATION FOR SEQ ID NO:517
20	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 17 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:517:
	Gly Arg Asp Tyr Tyr Asp Ser Gly Gly Tyr Phe Thr Val Ala Phe Asp 1 5 10 15
30	Ile
	(518) INFORMATION FOR SEQ ID NO:518
35	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 11 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
40	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:518:
<b>4</b> 5	Arg Ala Ser Gln Ser Ile Ser Arg Trp Leu Ala 1 5 10
₩.	(519) INFORMATION FOR SEQ ID NO:519
50	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 7 amino acids  (B) TYPE: amino acid  (C) TOPOLOGY: linear
	(ii) MOLECULE TYPE: peptide
EE	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:519:

	Glu Ala Ser Asn Asp Leu Ala 1 5
5	(520) INFORMATION FOR SEQ ID NO:520
10	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 5 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:520:
15	Asp Phe Tyr Met Glu 1 5
	(521) INFORMATION FOR SEQ ID NO:521
20	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 9 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
25	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:521:
30	Ile Ile Trp Asp Asp Gly Ser Asp Gln 1 5
	(522) INFORMATION FOR SEQ ID NO:522
15	<ul><li>(i) SEQUENCE CHARACTERISTICS:</li><li>(A) LENGTH: 11 amino acids</li><li>(B) TYPE: amino acid</li><li>(C) TOPOLOGY: linear</li></ul>
O	(ii) MOLECULE TYPE: peptide
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:522:
5	Gln Ala Ser Gln Ser Ile Ile Lys Tyr Leu Asn 1 5 10

Claims

 A method for determining how to humanize a rodent antibody or fragment thereof by resurfacing, said method comprising:

(a) determining the conformational structure of the variable region of said rodent antibody or fragment thereof by constructing a three-dimensional model of said rodent antibody variable region;

(b) generating sequence alignments from relative accessibility distributions from x-ray crystallographic structures of a sufficient number of rodent antibody variable region heavy and light chains to giv a set of heavy and light chain framework positions wherein said set is identical in 98% of said sufficient number of rodent antibody heavy and light chains;

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- (c) defining for said rodent antibody or fragment thereof to be humanized a set of heavy and light chain surface exposed amino acid residues using said set of framework positions generated in said step (b); (d) identifying from human antibody amino acid sequences a set of heavy and light chain surface exposed amino acid residues that is most closely identical to said set of surface exposed amino acid residues defined in said step (c), wherein said heavy and light chain from said human antibody are or are not naturally paired;
- (e) substituting, in the amino acid sequence of said rodent antibody or fragment thereof to be humanized said set of heavy and light chain surface exposed amino acid residues defined in said step (c) with said set of heavy and light chain surface exposed amino acid residues identified in said step (d);
- (f) constructing a three-dimensional model of said variable region of said rodent antibody or fragment thereof resulting from the substituting specified in said step (e);
- (g) identifying, by comparing said three-dimensional models constructed in said steps (a) and (f), any amino acid residues from said set identified in said step (d), that are within 5 Angstroms of any atom of any residue of the complementarity determining regions of said rodent antibody or fragment thereof to be humanized; and
- (h) changing any residues identified in said step (g) from the human to the original rodent amino acid residue to thereby define a rodent antibody humanizing set of surface exposed amino acid residues; with the proviso that said step (a) need not be conducted first, but must be conducted prior to said step (g).
- 2. The method of claim 1, wherein said rodent antibody is an antibody fragment.
- 3. The method of claim 2, wherein said rodent antibody fragment is a single chain antibody, a F<sub>V</sub> fragment, a Fab fragment, a Fab<sub>2</sub> fragment or a Fab' fragment.
- 4. The method of claim 1 or 2, wherein said step (d) identifies a set of naturally paired heavy and light chain surface exposed amino acid residues that is most closely identical to said set of surface exposed amino acid residues defined in said step (c).
- 5. The method of claim 1 or 2, wherein said surface exposed amino acid residues are those residues whose solvent accessibility is above 30%.
  - 6. The method of claim 1 or 2, wherein the rodent antibody or fragment thereof to be humanized is a murine antibody.
- 7. The method of claim 6, wherein said set of framework positions of surface exposed amino acid residu s is defined by the set shown in Table 1 and the alignments set forth in Figures 3A and 3B.

	Light Chain		
40 F	Position	Human	Mouse
1	1	D 51 E 34 A 5 S 5	D 76 Q 9 E 6
3	3	V 38 Q 24 S 24 Y 6	V 63 Q 22 L 5
45	5	T 61 L 37	Т 87
g	Ð	P 26 S 26 G 17 A 14 L 7	S 36 A 29 L 17 P 5
1	15	P 62 V 25 L 12	L47 P 30 V 8 A 7
50 1	18	R 57 S 18 T 13 P 6	R 38 K 22 S 13 Q 12 T 9
4	16	P 94	P 82 S 9
4	17	G 89	G 71 D 18
55 5	51	K 43 R 31	K 70 Q 13 R 8 T 5
6	<b>3</b> 3	G 91	G 98

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		1	1	
		66	D 43 S 25 A 9	D 38 A 26 S 26
	٠.	73 .	S 96	S 90 I 5
5		76	D 43 T 18 S 16 E 15	D 67 S 15 A 5 K 5
		86	P 44 A 27 S 17 T 8	A 50 P 11 T 8 E 7 Q 6
		87	E 71 D 11 G 7	E 91 D 6
10		111	K 74 R 12 N 6	K 93
		115	K 54 L 40	K 87 L 5
		116	R 60 G 33 S 5	R 89 G 9
15	***	117	Q 50 T 37 E 6 P 6	A74 Q 14 P 5 R 5
			Heavy Chain	
-	* *	Position	Human	Mouse
20		118	E 47 Q 46	E 59 Q 29 D 10
		120	Q 83 T 7	Q 68 K 26
		122	V 59 L 15 Q 13	Q 57 V 27 L 5 K 5
25		126	G 54 A 23 P 18	G 36 P 30 A 29
		127	G 53 E 22 A 14 D 7	E 45 G 43 S 6
		128	L 61 V 31 F 7	L 96
30		130	K 46 Q 41 E 5	K 52 Q 27 R 17
		131	P 95	P 91 A 5
		132	G 74 S 16 T 7	G 82 S 17
35		136	R 53 K 23 S 17 T 7	K 66 S 17 R 13
		143	G 96	G 98
40		145	T 46 S 32 N 9 I 7	T 63 S 19 N 7 A 5 D 5
		160	P 84 S 10	P 89 H 7
		161	G 93	G 71 E 24
45		162	K 76 Q 10 R 8	K 50 Q 30 N 10 H 5
		183	D 26 P 25 A 17 Q 10 T 7	E 31 P 22 D 17 A 12 Q 11
		184	S 70 K 9 P 8	K 42 S 37 T 6
50		186	K 53 Q 22 R 7 N 5	K 83 Q 7
		187	G 66 S 21 T 5	G 62 S 18 D 10
		195	T 30 D 26 N 19 K 7	T 36 K 30 N 26 D 6
55		196	S 91	S 76 A 16
		197	K 65 I 8 T 8 R 5	S 46 K 34 Q 11
		ı	1	]

	208	R 46 T 18 K 17 D 6	S 67 A 14 T 11			
.	209	A 50 P 21 S 13 T 8	E 88 D 7			
	210	E 46 A 18 D 13 S 9 Z 8 V 5	T 53 S 43			
L	212	Т 91				
	Table 1					

10

- 8. The method of claim 1 or 2, wherein the rodent antibody or fragment thereof to be humanized is murine antibody anti-N901.
- 9. The method of claim 8, wherein said set of framework positions of surface exposed amino acid residues is defined by the set shown in Table 1 and the alignments set forth in Figures 3A and 3B.

	Light Chain		
	Position	Human	Mouse
20	1	D 51 E 34 A 5 S 5	D76Q9E6
	3	V 38 Q 24 S 24 Y 6	V 63.Q 22 L 5
25	5	T 61 L 37	T 87
	9	P 26 S 26 G 17 A 14 L 7	S 36 A 29 L 17 P 5
	15	P 62 V 25 L 12	L 47 P 30 V 8 A 7
30	18	R 57 S 18 T 13 P 6	R 38 K 22 S 13 Q 12 T 9
30	46	P 94	P 82 S 9
	47	G 89	G 71 D 18
35	51	K 43 R 31	K 70 Q 13 R 8 T 5
33	63	G 91	G 98
	66	D 43 S 25 A 9	D 38 A 26 S 26
40	73	S 96	S 90 I 5
40	76	D 43 T 18 S 16 E 15	D 67 S 15 A 5 K 5
	86	P 44 A 27 S 17 T 8	A 50 P 11 T 8 E 7 Q 6
45	87	E 71 D 11 G 7	E 91 D 6
40	111	K 74 R 12 N 6	К 93
	115	K 54 L 40	K 87 L 5
50	116	R 60 G 33 S 5	R 89 G 9
<b></b>	117	Q 50 T 37 E 6 P 6	A 74 Q 14 P 5 R 5
	Heavy Chain		· .
55	Position	Human	Mous
JJ	118	E 47 Q 46	E 59 Q 29 D 10

	120	Q 83 T 7	Q 68 K 26	
	122	V 59 L 15 Q 13	Q 57 V 27 L 5 K 5	
5	126	G 54 A 23 P 18	G 36 P 30 A 29	
	127	G 53 E 22 A 14 D 7	E 45 G 43 S 6	
	128	L 61 V 31 F 7	L 96	
10	130	K 46 Q 41 E 5	K 52 Q 27 R 17	
	131	P 95	P 91 A 5	
	132	G 74 S 16 T 7	G 82 S 17	
<b>15</b>	136	R 53 K 23 S 17 T 7	K 66 S 17 R 13	
	143	G 96	G 98	
20	145	T 46 S 32 N 9 I 7	T 63 S 19 N 7 A 5 D 5	
20	160	P 84 S 10	P 89 H 7	
	161	G 93	G 71 E 24	
25	162	K 76 Q 10 R 8	K 50 Q 30 N 10 H 5	
	183	D 26 P 25 A 17 Q 10 T 7	E 31 P 22 D 17 A 12 Q 11	
	184	S 70 K 9 P 8	K 42 S 37 T 6	
30	186	K 53 Q 22 R 7 N 5	K 83 Q 7	
	187	G 66 S 21 T 5	G 62 S 18 D 10	
	195	T 30 D 26 N 19 K 7	T 36 K 30 N 26 D 6	
35	196	S 91	S 76 A 16	
	197	K 65 I 8 T 8 R 5	S 46 K 34 Q 11	
	208	R 46 T 18 K 17 D 6	S 67 A 14 T 11	
40	209	A 50 P 21 S 13 T 8	E 88 D 7	
	210	E 46 A 18 D 13 S 9 Z 8 V 5	T 53 S 43	
45	212	Т 91		
	Table 1			

- 10. A method for producing a humanized rodent antibody or fragment thereof from a rodent antibody or fragment thereof by resurfacing, said method comprising.
  - (I) carrying out the method of claim 1; and
  - (II) modifying the rodent antibody or fragment thereof by replacing the set of rodent antibody surface exposed amino acid residues with the rodent antibody humanizing set of surface exposed amino acid residues defined in said step (h).
  - 11. The method of claim 10, wherein said rodent antibody is an antibody fragment.
  - 12. The method of claim 11, wher in said rod int antibody fragment is a single chain antibody, a F<sub>V</sub> fragment,

- a Fab fragment, a Fab<sub>2</sub> fragment or a Fab' fragment.
- 13. The method of claim 10 or 11, wherein said step (d) identifies a set of naturally paired heavy and light chain surface exposed amino acid residues that is most closely identical to said set of surface exposed amino acid residues defined in said step (c).
- 14. The method of claim 10 or 11, wherein said surface exposed amino acid residues are those residues whose solvent accessibility is above 30%.
- 15. The method of claim 10 or 11, wherein the rodent antibody or fragment thereof to be humanized is a murin antibody.
  - 16. The method of claim 15, wherein said set of framework positions of surface exposed amino acid residues is defined by the set shown in Table 1 and the alignments set forth in Figures 3A and 3B.

15	Light Chain		
	Position	Human	Mouse
	1	D 51 E 34 A 5 S 5	D 76 Q 9 E 6
20	3	V 38 Q 24 S 24 Y 6	V 63 Q 22 L 5
	5	T 61 L 37	T 87
	9	P 26 S 26 G 17 A 14 L 7	S 36 A 29 L 17 P 5
25	15	P 62 V 25 L 12	L 47 P 30 V 8 A 7
	18	R 57 S 18 T 13 P 6	R 38 K 22 S 13 Q 12 T 9
	46	P 94	P 82 S 9
30	47	G 89	G 71 D 18
	51	K 43 R 31	K 70 Q 13 R 8 T 5
	63	G 91	G 98
35	66	D 43 S 25 A 9	D 38 A 26 S 26
	73	S 96	S 90 I 5
	76	D 43 T 18 S 16 E 15	D 67 S 15 A 5 K 5
	86	P 44 A 27 S 17 T 8	A 50 P 11 T 8 E 7 Q 6
	87	E 71 D 11 G 7	E 91 D 6
	111	K 74 R 12 N 6	K 93
45	115	K 54 L 40	K 87 L 5
	116	R 60 G 33 S 5	R 89 G 9
	117	Q 50 T 37 E 6 P 6	A 74 Q 14 P 5 R 5
50	Heavy Chain		
	Position	Human	Mouse
	118	E 47 Q 46	E 59 Q 29 D 10
55	120	Q 83 T 7	Q 68 K 26
	122	V 59 L 15 Q 13	Q 57 V 27 L 5 K 5

	126	G 54 A 23 P 18	G 36 P 30 A 29
•	127	G 53 E 22 A 14 D 7	E 45 G 43 S 6
5	128	L 61 V 31 F 7	L 96
	130	K 46 Q 41 E 5	K 52 Q 27 R 17
	131	P 95	P 91 A 5
10	132	G 74 S 16 T 7	G 82 S 17
	136	R 53 K 23 S 17 T 7	K 66 S 17 R 13
	143	G 96	G 98
15	145	T 46 S 32 N 9 I 7	T 63 S 19 N 7 A 5 D 5
	160	P 84 S 10	P 89 H 7
20	161	G 93	G 71 E 24
	162 .	K 76 Q 10 R 8	K 50 Q 30 N 10 H 5
	183	D 26 P 25 A 17 Q 10 T 7	E 31 P 22 D 17 A 12 Q 11
25	184	S 70 K 9 P 8	K 42 S 37 T 6
	186	K 53 Q 22 R 7 N 5	K 83 Q 7
	187	G 66 S 21 T 5	G 62 S 18 D 10
30	195	T 30 D 26 N 19 K 7	T 36 K 30 N 26 D 6
•	196	S 91	S 76 A 16
	197	K 65   8 T 8 R 5	S 46 K 34 Q 11
35	208	R 46 T 18 K 17 D 6	S 67 A 14 T 11
	209	A 50 P 21 S 13 T 8	E 88 D 7
	210	E 46 A 18 D 13 S 9 Z 8 V 5	T 53 S 43
40	212	Т 91	
		Table 1	

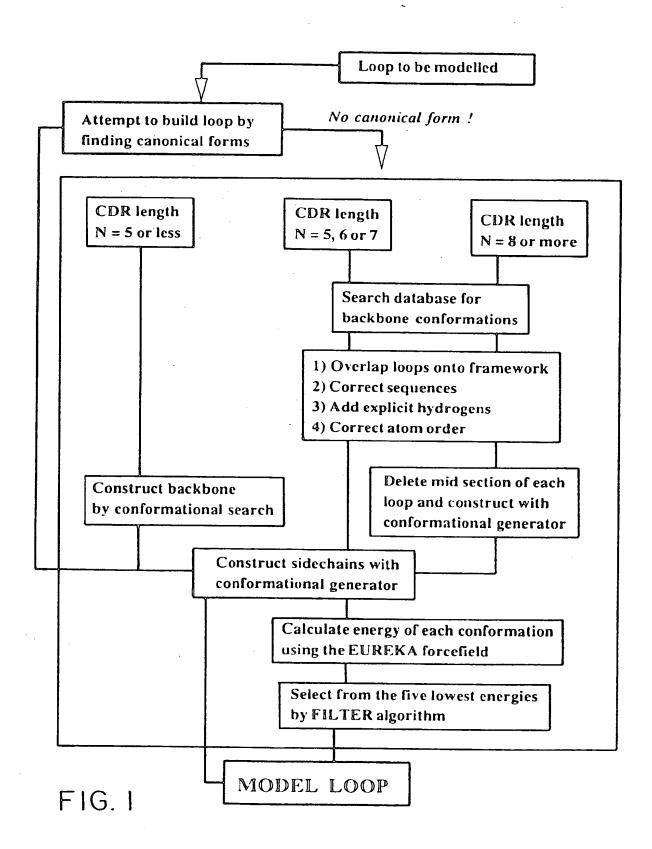
- 45 17. The method of claim 10 or 11, wherein the rodent antibody or fragment thereof to be humanized is murin antibody anti-N901.
  - 18. The method of claim 17, wherein said set of framework positions of surface exposed amino acid residues is defined by the set shown in Table 1 and the alignments set forth in Figures 3A and 3B.

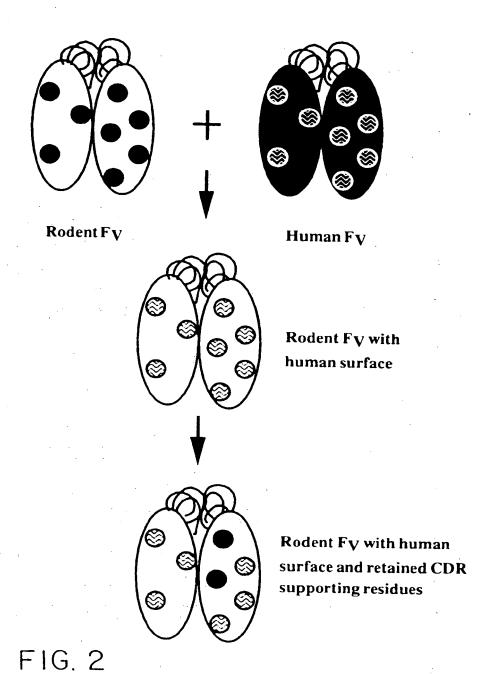
Light Chain			
Position	Human	Mous	
1	D 51 E 34 A 5 S 5	D 76 Q 9 E 6	
3	V 38 Q 24 S 24 Y 6	V 63 Q 22 L 5	

50

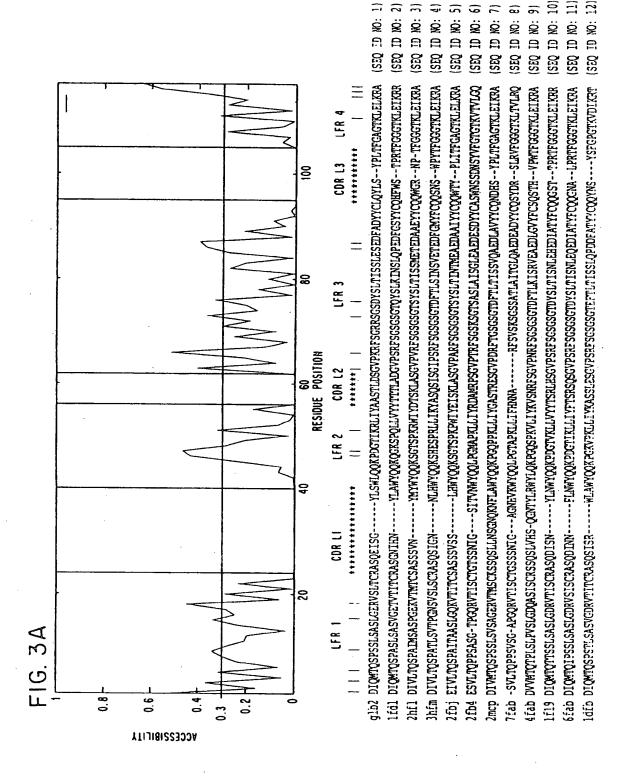
	5	T 61 L 37	T 87
	9	P 26 S 26 G 17 A 14 L 7	S 36 A 29 L 17 P 5
5	15	P 62 V 25 L 12	L 47 P 30 V 8 A 7
	18	R 57 S 18 T 13 P 6	R 38 K 22 S 13 Q 12 T 9
	46	P 94	P 82 S 9
10	47	G 89	G 71 D 18
	51	K 43 R 31	K 70 Q 13 R 8 T 5
	63	G 91	G 98
15	66	D 43 S 25 A 9	D 38 A 26 S 26
	73	S 96	S 90 I 5
	76	D 43 T 18 S 16 E 15	D 67 S 15 A 5 K 5
20	86	P 44 A 27 S 17 T 8	A 50 P 11 T 8 E 7 Q 6
	87	E 71 D 11 G 7	E 91 D 6
	111	K 74 R 12 N 6	К 93
25	115	K 54 L 40	K 87 L 5
	116	R 60 G 33 S 5	R 89 G 9
	117	Q 50 T 37 E 6 P 6	A 74 Q 14 P 5 R 5
30	Heavy Chain		
	Position	Human	Mouse
	Position 118	Human E 47 Q 46	Mouse E 59 Q 29 D 10
35			
35	118	E 47 Q 46	E 59 Q 29 D 10
35	118 120	E 47 Q 46 Q 83 T 7	E 59 Q 29 D 10 Q 68 K 26
35	118 120 122	E 47 Q 46 Q 83 T 7 V 59 L 15 Q 13	E 59 Q 29 D 10 Q 68 K 26 Q 57 V 27 L 5 K 5
	118 120 122 126	E 47 Q 46 Q 83 T 7 V 59 L 15 Q 13 G 54 A 23 P 18	E 59 Q 29 D 10 Q 68 K 26 Q 57 V 27 L 5 K 5 G 36 P 30 A 29
	118 120 122 126 127	E 47 Q 46  Q 83 T 7  V 59 L 15 Q 13  G 54 A 23 P 18  G 53 E 22 A 14 D 7	E 59 Q 29 D 10 Q 68 K 26 Q 57 V 27 L 5 K 5 G 36 P 30 A 29 E 45 G 43 S 6
	118 120 122 126 127 128	E 47 Q 46  Q 83 T 7  V 59 L 15 Q 13  G 54 A 23 P 18  G 53 E 22 A 14 D 7  L 61 V 31 F 7	E 59 Q 29 D 10  Q 68 K 26  Q 57 V 27 L 5 K 5  G 36 P 30 A 29  E 45 G 43 S 6  L 96
<b>40</b>	118 120 122 126 127 128 130	E 47 Q 46  Q 83 T 7  V 59 L 15 Q 13  G 54 A 23 P 18  G 53 E 22 A 14 D 7  L 61 V 31 F 7  K 46 Q 41 E 5	E 59 Q 29 D 10  Q 68 K 26  Q 57 V 27 L 5 K 5  G 36 P 30 A 29  E 45 G 43 S 6  L 96  K 52 Q 27 R 17
40	118 120 122 126 127 128 130	E 47 Q 46  Q 83 T 7  V 59 L 15 Q 13  G 54 A 23 P 18  G 53 E 22 A 14 D 7  L 61 V 31 F 7  K 46 Q 41 E 5  P 95	E 59 Q 29 D 10  Q 68 K 26  Q 57 V 27 L 5 K 5  G 36 P 30 A 29  E 45 G 43 S 6  L 96  K 52 Q 27 R 17  P 91 A 5
<b>40</b>	118 120 122 126 127 128 130 131	E 47 Q 46  Q 83 T 7  V 59 L 15 Q 13  G 54 A 23 P 18  G 53 E 22 A 14 D 7  L 61 V 31 F 7  K 46 Q 41 E 5  P 95  G 74 S 16 T 7	E 59 Q 29 D 10  Q 68 K 26  Q 57 V 27 L 5 K 5  G 36 P 30 A 29  E 45 G 43 S 6  L 96  K 52 Q 27 R 17  P 91 A 5  G 82 S 17
40	118 120 122 126 127 128 130 131 132	E 47 Q 46  Q 83 T 7  V 59 L 15 Q 13  G 54 A 23 P 18  G 53 E 22 A 14 D 7  L 61 V 31 F 7  K 46 Q 41 E 5  P 95  G 74 S 16 T 7  R 53 K 23 S 17 T 7	E 59 Q 29 D 10  Q 68 K 26  Q 57 V 27 L 5 K 5  G 36 P 30 A 29  E 45 G 43 S 6  L 96  K 52 Q 27 R 17  P 91 A 5  G 82 S 17  K 66 S 17 R 13  G 98
40 45 50	118 120 122 126 127 128 130 131 132 136	E 47 Q 46  Q 83 T 7  V 59 L 15 Q 13  G 54 A 23 P 18  G 53 E 22 A 14 D 7  L 61 V 31 F 7  K 46 Q 41 E 5  P 95  G 74 S 16 T 7  R 53 K 23 S 17 T 7  G 96  T 46 S 32 N 9 I 7	E 59 Q 29 D 10  Q 68 K 26  Q 57 V 27 L 5 K 5  G 36 P 30 A 29  E 45 G 43 S 6  L 96  K 52 Q 27 R 17  P 91 A 5  G 82 S 17  K 66 S 17 R 13  G 98  T 63 S 19 N 7 A 5 D 5
45	118 120 122 126 127 128 130 131 132 136 143	E 47 Q 46  Q 83 T 7  V 59 L 15 Q 13  G 54 A 23 P 18  G 53 E 22 A 14 D 7  L 61 V 31 F 7  K 46 Q 41 E 5  P 95  G 74 S 16 T 7  R 53 K 23 S 17 T 7  G 96	E 59 Q 29 D 10  Q 68 K 26  Q 57 V 27 L 5 K 5  G 36 P 30 A 29  E 45 G 43 S 6  L 96  K 52 Q 27 R 17  P 91 A 5  G 82 S 17  K 66 S 17 R 13  G 98

	162	K 76 Q 10 R 8	K 50 Q 30 N 10 H 5	
	183	D 26 P 25 A 17 Q 10 T 7	E 31 P 22 D 17 A 12 Q 11	
5	184	S 70 K 9 P 8	K 42 S 37 T 6	
	186	K 53 Q 22 R 7 N 5	K 83 Q 7	
	187	G 66 S 21 T 5	G 62 S 18 D 10	
10	195	T 30 D 26 N 19 K 7	T 36 K 30 N 26 D 6	
	196	S 91	S 76 A 16	
	197	K 65 I 8 T 8 R 5	S 46 K 34 Q 11	
15	208	R 46 T 18 K 17 D 6	S 67 A 14 T 11	
	209	A 50 P 21 S 13 T 8	E 88 D 7	
20	210	E 46 A 18 D 13 S 9 Z 8 V 5	T 53 S 43	
20	212	T 91		
		Table 1		





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14)

16)

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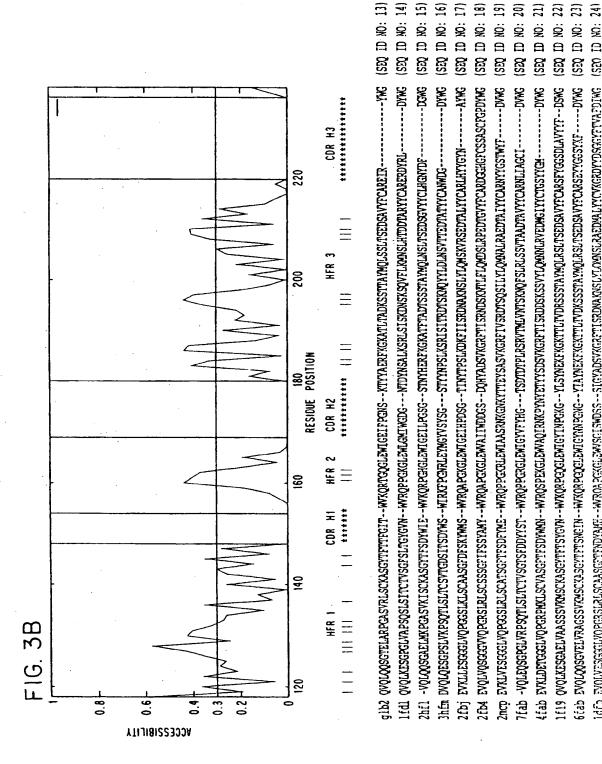
18) 19) 20)

21)

22) 23) 24)

(330

JVOLVESAGALVOPGASLALSARARGFISVDYAMG - "WROA PGKGLJWYSGISWDSS --SIGYADSYKGARTISRDNAKUSLYLANNSGAAEDMALYYCVKGADYYDSAGYFTVAPDIWG



## FIG. 4A Light Chain Sequences

70	VPDRFSG	VPDRFSG	VPDRFSG	VPDRFSG	VPDRFSG	VPDRFSG	/PDRFSG
09	KVSNRFS G	RDAMRPS G	KVSNRFS G	KVSNRDS G	KVSNRFS G	WASTRES G	KVSNRFS GV L2
20	WFLQKPGQSPKLLIY	:QSVLTQPPSASG-TPGQRVTISC SGTSSNIGSSTVN WYQQLPGMAPKLLIY RDAMRPS GVPDRFSG	QVLMTQTPSSLPVTLGQQASISC RSSQIIIHSDGNTY-LE WFLQKPGQSPKLLIY KVSNRFS GVPDRFSG	:DVVMTQSPLSLPVTLGQPASISC RSSQSLVYSDGNTY-LN WFQQRPGQSPRRLIY KVSNRDS GVPDRFSG	:DVLMTQSPLSLPVTLGQPASISC RSSQIIIHSDGNTY-LE WFQQRPGQSPRLLIY KVSNRFS GVPDRFSG	:DIVMTQSPDSLAVSLGERATINC KSSQSVLYSSNNKNYLA WYQQKPGQPPKLLIY WASTRES GVPDRFSG	: DVLMTQTPDSLPVSLGDRASISC RSSQIIIHSDGNTY-LE WFLQKPGQSPKLLIY KVSNRFS GVPDRFSG [ L1 ]
	QIIIHSDGNTY-LE	VIGSSTVN	IIHSDGNTY-LE	.VYSDGNTY-LN	I HSDGNTY-LE	/LYSSNNKNYLA	[IHSDGNTY-LE L1
	RSSQI	SGTSSI	RSSQI	RSSQSI	RSSQI	: KSSQS1	RSSQII
20	PVSLGDQASISC	3-TPGQRVTISC	PVŤĽGQQASISC	PVTLGQPASISC	PVTLGQPASISC	AVSLGERATINC	PVSLGDRASISC
10	DVLMTQTPLSL	:QSVLTQPPSAS(	QVLMTQTPSSL	DVVMTQSPLSL	: DVLMTQSPLSLI	:DIVMTQSPDSLA	:DVLMTQTPDSL
				( מפיג		Surf	
	1 N901L	2 KOL	3 N901L/KOL	4 KV2F\$HUMAN [most identica]	5 N901L/KV2F	6 KV4B\$HUMAN [most identica]	7 N901L/KV4B [Resurfaced]

	08	06	100				
N901L	SGSGTDFTLMISRVEAEDLGVYYC FQGSHVPHT FGGGTKLEI-	EAEDLGVYYC	EDLGVYYC FOGSHVPHT	FGGGTKLEI-		(SEQ ID NO: 25)	25)
KOL	SKSGASASLAIGGLQSEDETDYYC AAWDVSLNAYV FGTGTKVTVL	QSEDETDYYC	AAWDVSLNAYV	FGTGTKVTVL	(44)	(SEQ ID NO: 26)	26)
N901L/KOL	SGSGTSFTLAISRVEAEDEGVYYC FQGSHVPHT FGGGTKLEI-	EAEDEGVYYC	FQGSHVPHT	FGGGTKLEI-	(104)	(SEQ ID NO: 27)	27)
KV2F\$HUMAN	:SGSGTDFTLKISRVEAEDVGVYYC MQGTHWSWT FGQGTKVEIK	EAEDVGVYYC	MQGTHWSWT	FGGGTKVEIK	( 81)	(SEQ ID NO: 28)	28)
_	SGSGTDFTLKISRVEAEDVGVYYC FQGSHVPHT FGGGTKVEI-	EAEDVGVYYC	FQGSHVPHT	FGGGTKVEI -	(101)	(SEQ ID NO: 29)	29)
KV4B\$HUMAN	SGSGTDFTLTISSLQAEDVAVYYC QQYDTIPT FGGGTKVEIK	QAEDVAVYYC	QQYDTIPT	FGGGTKVEIK	(71)	(SEQ ID NO: 30)	30)
(most identical sull) N901L/KV4B [Resurfaced]	SGSGTDFTLMISRVEAEDLGVYYC FQGSHVPHT FGGGTKLEI-	EAEDLGVYYC	FQGSHVPHT	FGGGTKLEI-	(109)	(SEQ ID NO: 31)	31)

FIG. 4B

					: 32)	33)	34)	35)	36)	37)	38)	
					ON CI	ON Q	ON O	D NO	Ö.	D NO:	D NO:	
VKG	VKG	VKG	VKG VKG		(SEQ. 1	SEQ I	SEQ I	SEQ I	SEQ I	SEQ I	SEQ I	
HADT	YADS       	YADS     YADS	YVDS'       HADS'						03) (	74) (	10) (	
+ TIY	DQH	NKY	EKY			<u> </u>	(1	~	C	_	(1)	
YISSGSF	IIWDDGS.	VISYDGS- YISSGSF-	NIKQDGS- YISSGSF-	240	GTTVTVS	GTPVTVS	GTTVTVS	GTLVTVS	GTLVTVS	! ! !	GTTVTVS	
LEWVA	LEWVA	LEWVA	EWVA EWVA		MGC	∑ .wgg	Y WGQ	Y WGQ	Y WGQ	1 .	Y WGQ	
WVRQAPEKG	avrqapgkgi   avrqapgkgi	AVRQA PGKGI   AVRQA PGKGI	VVRQA PGKGI I VVRQA PGKGI	230	0J	SSASCFGPD	! !	;	 	1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	1	
		SYAMH I	SYWMS I SFGMH V H1 ]	220	MRKGYAN	DGGHGFC	MRKGYAN	DRKDWGM	MRKGYAM		MRKGYAM [	
		RLSCAASGFTFS (	RLSCAASGFTFS S	210	SLRSEDTAMYYCAF	SLRPEDTGVYFCAR	SLRSEDTAMYYCAR	SLRAEDTAVYYCAR	SLRAEDTAVYYCAR	SLRAEDTAVYYCAR	Slraedtamyycar	
GGGLVQPGGSF	GGGVVQPGRSI 	GGGVVQPGRSI 	GGGLVQPGGSL   GGGLVQPGGSL	200	npkntlflomt	DSKNTLFLQMD	DPKNTLFLOMT	NSKNTLYLQMN	NSKNTLYLOMN	NAKNSLYLQMN	VAKNTLFLQMT	
: DVQLVES	: EVQLVQS   : EVQLVES	:QVQLVES   :QVQLVES	: EVQLV ! : EVQLV	190		: RFTI SRN	RFTISRD	RFTISRD	RFTISRD	RFTISRD	RFTISRD	
		sed]	surf					sed	•	surf		
1 N901H	2 KOL 3 N901H/KOL	G36005 [most identical N901H/G36005 [CDR grafted]			1 N901H	2 KOL	3 N901H/KOL			6 PL0123 [most identical	7 N901H/PL0123 [Resurfaced]	
		N901H  SDVQLVESGGGLVQPGGSRKLSCAASGFTFS SFGMH  KOL  SEVQLVQSGGGVVQPGRSLRLSCSSSGFIFS SYAMY  1	N901H  KOL  EVQLVQSGGGLVQPGGSRKLSCAASGFTFS SFGMH  KOL  EVQLVQSGGGVVQPGRSLRLSCSSGFIFS SYAMY  1	KOL  :EVQLVQSGGGLVQPGGSRKLSCAASGFTFS SFGMH	KOL : EVQLVQSGGGVVQPGGSRKLSCAASGFTFS SFGMH	: EVQLVESGGGLVQPGGSRKLSCAASGFTFS SFGMH : EVQLVQSGGGVVQPGRSLRLSCAASGFIFS SYAMY i dentical seq]	11H/KOL  EVQLVGSGGGLVQPGGSRKLSCAASGFTFS SFGMH WVRQAPEKGLEWVA IIWDDGSTIY HADTVKG  EVQLVGSGGGVVQPGRSLRLSCAASGFTFS SFGMH WVRQAPGKGLEWVA IIWDDGSDQH YADSVKG  11H/KOL  EVQLVESGGGVVQPGRSLRLSCAASGFTFS SFGMH WVRQAPGKGLEWVA VISYDGSTIY HADSVKG  12t identical seq!   VVQLVESGGGVVQPGRSLRLSCAASGFTFS SFGMH WVRQAPGKGLEWVA VISYDGSNKY YADSVKG  12 identical seq!   VVQLVESGGGVVQPGRSLRLSCAASGFTFS SFGMH WVRQAPGKGLEWVA VISYDGSTIY YADSVKG  12 identical surf!   VVQLVESGGGLVQPGGSLRLSCAASGFTFS SFGMH WVRQAPGKGLEWVA VISSGSFTIY YADSVKG  12 identical surf!   VVQLVESGGGLVQPGGSLRLSCAASGFTFS SFGMH WVRQAPGKGLEWVA VISSGSFTIY HADSVKG  12 identical surf!   VVQLVESGGGLVQPGGSLRLSCAASGFTFS SFGMH WVRQAPGKGLEWVA VISSGSFTIY HADSVKG  14 ill   VVGLVESGGGLVQPGGSLRLSCAASGFTFS SFGMH WVRQAPGKGLEWVA VISSGSFTIY HADSVKG  14 ill   STTISRNDFKNTLFLQMTSLRSEDTAMYYCAR MRKGYAMDY WGQGTFVTVS (SEQ ID NO:  15   VVGLVESGGGLVQPGSRLRSEDTAMYYCAR MRKGYAMDY WGQGTFVTVS (77) (SEQ ID NO:  16   VVGLVESGGGLVQPGSRLRSEDTAMYYCAR DGGHGFCSSASCFGDDY WGQGTFVTVS (77) (SEQ ID NO:  17   SETISRNDSKNTLFLQMTSLRSEDTAMYYCAR DGGHGFCSSASCFGDDY WGQGTFVTVS (77) (SEQ ID NO:  18   VVGLVESGGLVQPGSRLRSEDTAMYYCAR DGGHGFCSSASCFGDDY WGQGTFVTVS (77) (SEQ ID NO:  19   VVGLVESGGGLVQPGSRLRSEDTAMYYCAR DGGHGFCSSASCFGDDY WGQGTFVTVS (77) (SEQ ID NO:  10   VVGLVESGGGLVQPGSRLRSEDTAMYYCAR DGGHGFCSSASCFGDDY WGQGTFVTVS (77) (SEQ ID NO:  10   VVGLVESGGGLVQPGSRLRSEDTAMYYCAR DGGHGFCSSASCFGDDY WGQGTFVTVS (77) (SEQ ID NO:  10   VVGLVESGGGLVQPGSRLRSEDTAMYYCAR DGGHGFCSSASCFGDDY WGQGTFVTVS (77) (SEQ ID NO:  11   VVGLVESGGGLVQPGSRLRSEDTAMYYCAR DGGHGFCSSASCFGDDY WGGGTFVTVS (77) (SEQ ID NO:  11   VVGLVESGGGLVQPGSRLRSEDTAMYYCAR DGGHGFCSSASCFGDDY WGGGTFVTVS (77) (SEQ ID NO:  12   VVGLVESGGGLVQPGSRLRSEDTAMYYCAR DGGHGFCSSASCFGDDY WGGGTFVTVS (77) (SEQ ID NO:  15   VVGLVESGGGLVQPGSRLRSEDTAMYYCAR DGGHGFCSSASCFGDDY WGGGTFVTVS (77) (SEQ ID NO:  15   VVGLVESGGGLVQPGSRLRSEDTAMYYCAR DGGHGFCSSASCFGDDY WGGGTFVTVS (77) (SEQ ID NO:  15   VVGLVESGGGLVQPGSRLRSEDTAMYYCAR DGGHGFCSSASCFGDDY WGGGTFVTVS (77) (V	11H/KOL  1 EVQLVGSGGGLVQPGGSRKLSCASGFTFS SFGMH WVRQAPEKGLEWVA YISSGSFTIY HADTVKG  1 EVQLVGSGGGVVQPGRSLRLSCASGFTFS SYAMY WVRQAPGKGLEWVA YISSGSFTIY HADSVKG  1 EVQLVESGGGVVQPGRSLRLSCASGFTFS SYAMH WVRQAPGKGLEWVA YISSGSFTIY HADSVKG  1 C C C C C C C C C C C C C C C C C C	:EVQLVQSGGGLVQPGGSRKLSCAASGFTFS SFGMH WYRQAPEKGLEWVA YISSGSFTIY HADD:	11   11   12   12   12   13   14   15   15   15   15   15   15   15	BVQLVESGGGLVQPGGSRKLSCASGFTES SFGHH WYRQAPEKGLEWVA YISSGSFTIY HADD   COULDESGGGVVQPGRSLRLSCSSGFIFS SYAMY WYRQAPEKGLEWVA YISSGSFTIY HADD   COULDESGGGVVQPGRSLRLSCAASGFIFS SYAMY WYRQAPGKGLEWVA YISSGFTIY HADD   COULDESGGGVVQPGRSLRLSCAASGFTFS SYAMH WYRQAPGKGLEWVA YISSGFTIY YADD   COULDESGGGVVQPGRSLRLSCAASGFTFS SYAMH WYRQAPGKGLEWVA YISSGSFTIY YADD   COULDESGGGVVQPGRSLRLSCAASGFTFS SFGMH WYRQAPGKGLEWVA YISSGSFTIY YADD   COULDESGGGVVQPGRSLRLSCAASGFTFS SFGMH WYRQAPGKGLEWVA YISSGSFTIY YADD   COULDESGGGLVQPGGSLRLSCAASGFTFS SFGMH WYRQAPGKGLEWVA YISSGSFTIY YADD   COULDESGGGLVQPGGSLRLSCAASGFTFS SFGMH WYRQAPGKGLEWVA YISSGSFTIY HADD   COULDESGGGLVQPGGSLRLSCAASGFTFS SFGMH WYRQAPGKGLEWVA YISSGSFTIY YADD   COULDESGGGLVQPGGSLRLSCAASGFTFS SFGMH WYRQAPGKGLEWVA YISSGSFTIY HADD   COULDESGGGLVQPGGSLRLSCAASGFTFS SFGMH WYRQAPGKGLEWVA YISSGSFTIY YADD   COULDESGGGLVQPGGSLRLSCAASGFTFS SFGMH WYRQAPGKGLEWVA YISSGSFTIY HADD   COULDESGGGLVQPGGSLRLSCAASGFTFS SFGMH WYRQAPGKGLEWVA YISSGSFTIY YADD   COULDESGGGLVQPGGSLRLSCAASGFTFS SFGMH WYRQAPGKGLEWVA YISSGSFTIY HADD   COULDESGGGLVQPGGSLRLSCAASGFTFS SFGMH WYRQAPGKGLEWVA YISSGSFTIY YADD   COULDESGGGLVQPGGSLRSCAASGFTFS SFGMH WYRQAPGKGLEWVA YISSGSFTIY YADD   COULDESGGGLVQPGGSLRSCAASGFTFS SFGMH WYRQAPGKGLEWV	19

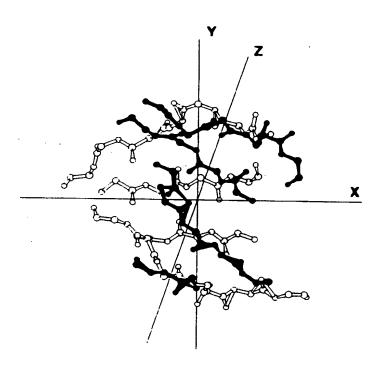
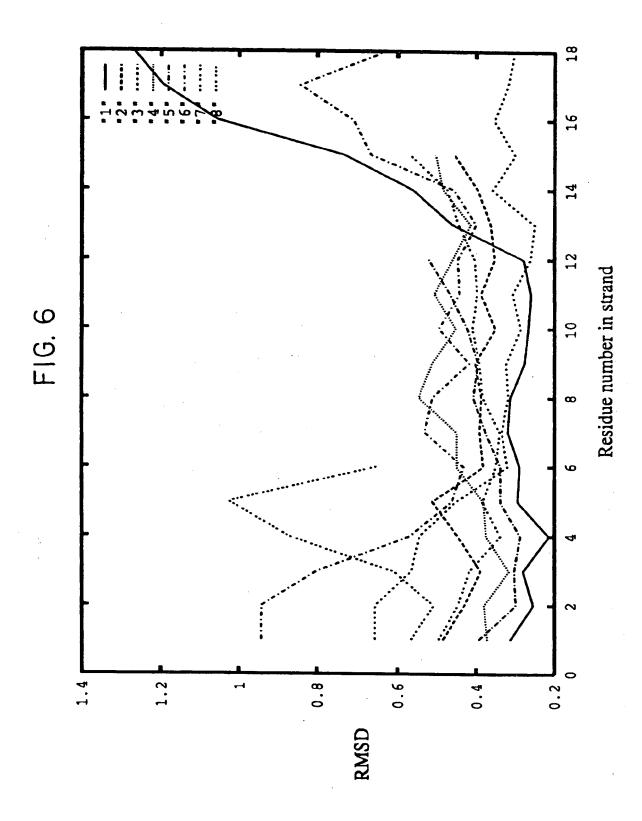
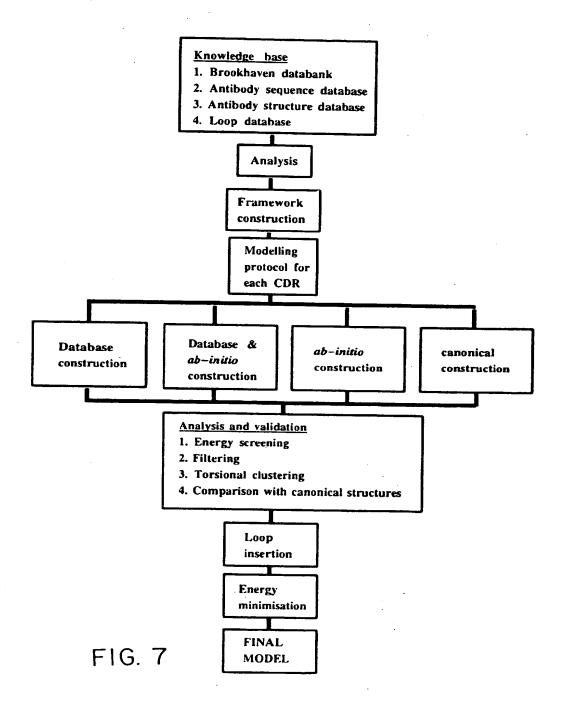
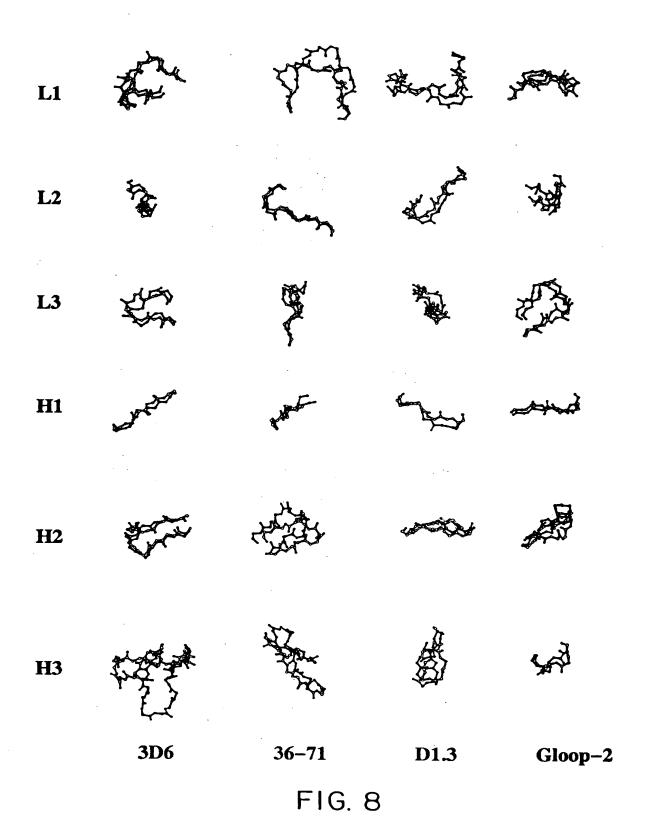
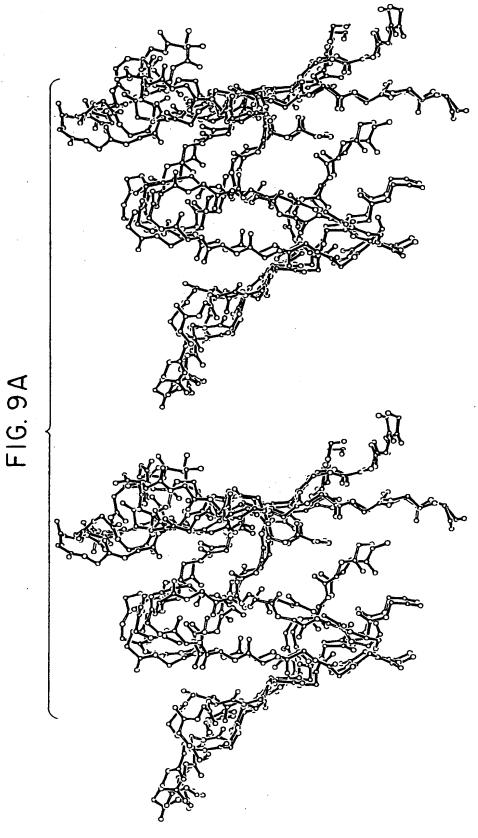


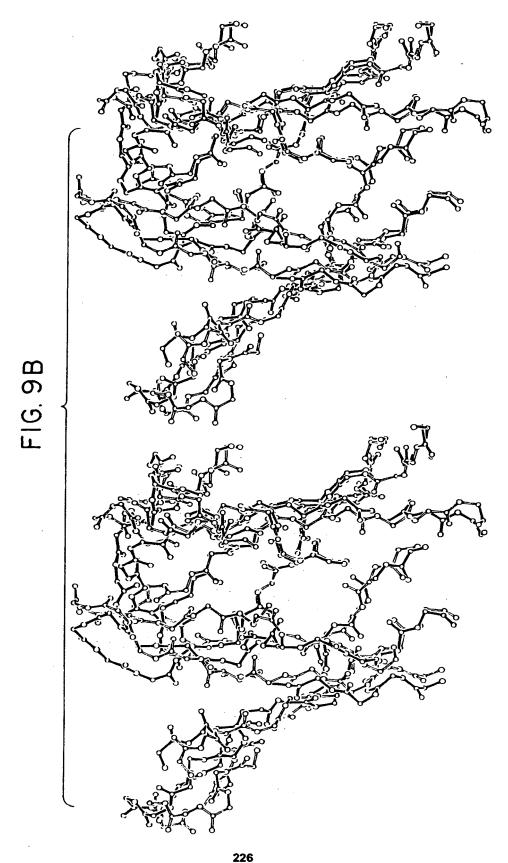
FIG. 5

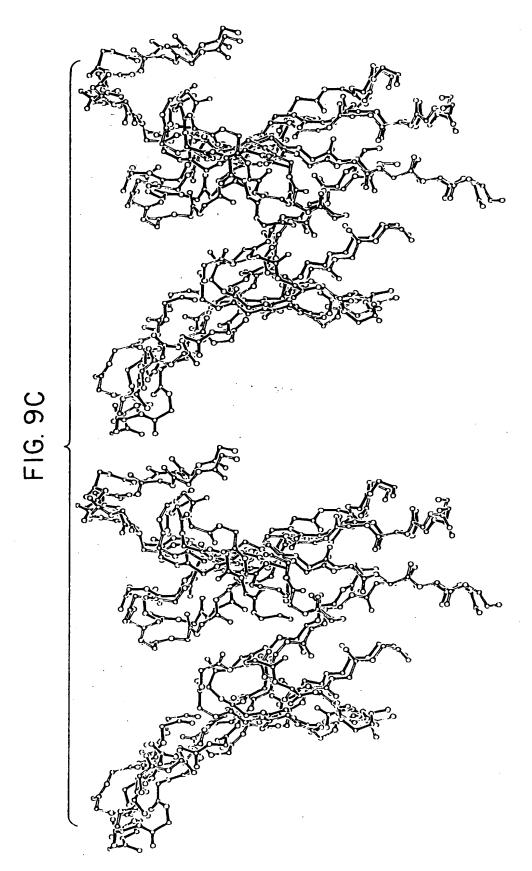


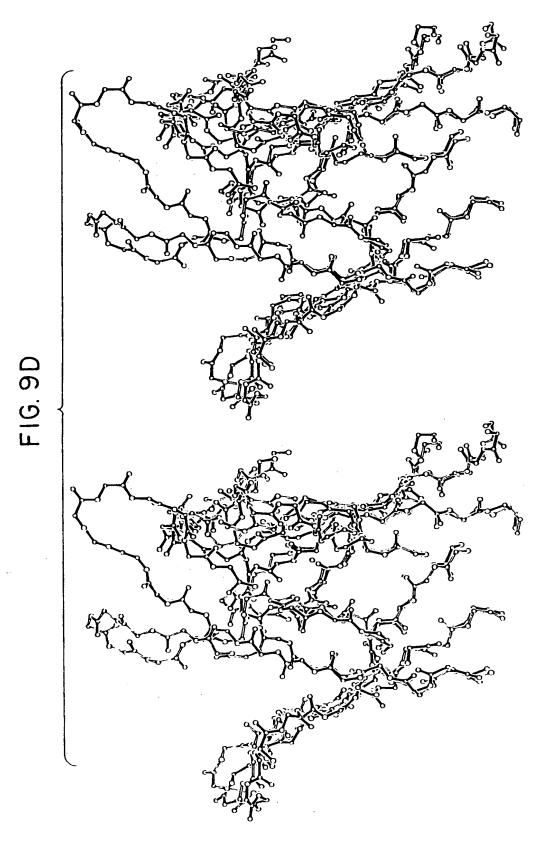












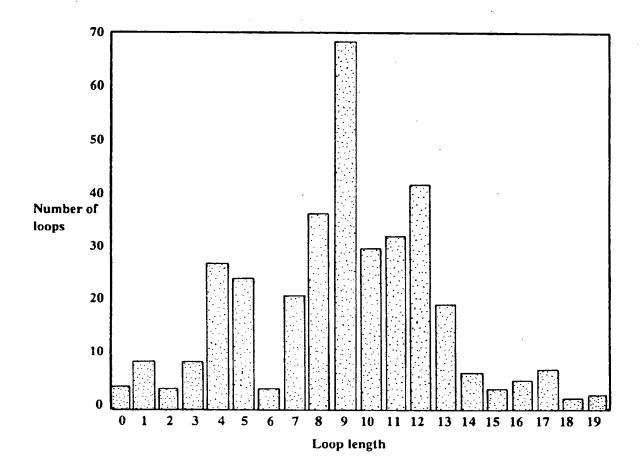


FIG. 10



## **EUROPEAN SEARCH REPORT**

Application Number

EP 93 30 7051

ategory	Citation of document with it of relevant pa	ndication, where appropriate,	Relevant to claim	CLASSIFICATION OF THE APPLICATION (Int. Cl.5)
D,A	MOLECULAR IMMUNOLO vol. 28, no. 4/5, 1 pages 489 - 498 PADLAN A E 'POSSIBL REDUCING THE IMMUNO	GY 991, GB E PROCEDURE FOR GENICITY OF ANTIBODY ILE PRESERVING THEIR ERTIES'		C12N15/13 C12N15/62 C07K15/00 C12P21/08
D, A	WO-A-9 109 967 (CEL 11 July 1991 * p. 5, second para paragraph, "Ration	-		
P,A	EP-A-0 519 596 (MER 23 December 1992 * Claims *	CK & CO. INC.)		
	•			
				TECHNICAL FIELDS SEARCHED (Int. Cl.5)
				C07K
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	The present search report has	been drawn up for all claims		
	Place of search	Date of completion of the search	<del>'.</del>	Exeminer
	MUNICH	12 JANUARY 1994		Germinario C.
Y:pa do A:te O:no	CATEGORY OF CITED DOCUME rticularly relevant if taken alone rticularly relevant if combined with an current of the same category chnological background in-written disclosure termediate document	E : earlier patent e ziter the filing nother D : document citer L : document citer	focument, but purdate I in the application of the reason	blished on, or